Table of Contents

14	CARDIOVASCU	LAR ASSESSMENT	14-1
1	4.1 INTRODUCT	ION	14-1
	14.1.1 Backgro	ound	14-1
		ry of Previous Analyses of the Air Force Health Study	
		Baseline Study Summary Results	
		Follow-up Study Summary Results	
		Follow-up Study Summary Results	
		m Dioxin Analysis of 1987 Follow-up Study Summary Results	
		Follow-up Study Summary Results	
		ters for the 1997 Cardiovascular Assessment	
		endent Variables	
		Medical Records Data	
	14.1.3.1.2	Physical Examination Data and Self-reported Questionnaire Data	14-5
	14.1.3.2 Cova	uriates	
		al Methods	
	14.1.4.1 Long	gitudinal Analysis	14-12
1	4.2 RESULTS	•	14-12
	14.2.1 Depend	ent Variable-Covariate Associations	14-12
		re Analysis	
		ical Records Variables	
	14.2.2.1.1	Essential Hypertension	
	14.2.2.1.2	Heart Disease (Excluding Essential Hypertension)	
	14.2.2.1.3	Myocardial Infarction	
	14.2.2.1.4	Stroke or Transient Ischemic Attack	14-25
	14.2.2.2 Phys	ical Examination Variables – Central Cardiac Function	14-28
	14.2.2.2.1	Systolic Blood Pressure (Continuous)	
	14.2.2.2.2	Systolic Blood Pressure (Discrete)	14-31
	14.2.2.2.3	Diastolic Blood Pressure (Continuous)	14-34
	14.2.2.2.4	Diastolic Blood Pressure (Discrete)	14-38
	14.2.2.2.5	Heart Sounds	14-40
	14.2.2.2.6	Overall Electrocardiograph	
	14.2.2.2.7	Right Bundle Branch Block	
	14.2.2.2.8	Left Bundle Branch Block	
	14.2.2.2.9	Non-Specific ST- and T-Wave Changes	
		Bradycardia	
	14.2.2.2.11	Tachycardia	
	14.2.2.2.12	Arrhythmia	
	14.2.2.2.13	Evidence of Prior Myocardial Infarction	
	14.2.2.2.14	ECG: Other Diagnoses	
		ical Examination Variables – Peripheral Vascular Function	
	14.2.2.3.1	Funduscopic Examination	
	14.2.2.3.2	Carotid Bruits	
	14.2.2.3.3	Radial Pulses	
	14.2.2.3.4	Femoral Pulses	
	14.2.2.3.5	Popliteal Pulses	
	14.2.2.3.6 14.2.2.3.7	Dorsalis Pedis Pulses Posterior Tibial Pulses	
	14.2.2.3.7	POSICIOU LIDIAL PHISCS	14-81

14.2.2.3.8	Leg Pulses	14-83				
14.2.2.3.9	Peripheral Pulses	14-85				
14.2.2.3.10	ICVÎ Index					
14.2.3 Longitu	ıdinal Analysis	14-90				
14.2.3.1 Phys	sical Examination Variables	14-91				
14.2.3.1.1	Systolic Blood Pressure (Continuous)	14-91				
14.2.3.1.2	Systolic Blood Pressure (Discrete)	14-94				
14.2.3.1.3	Femoral Pulses	14-97				
14.2.3.1.4	Popliteal Pulses	14-100				
14.2.3.1.5	Dorsalis Pedis Pulses	14-103				
14.2.3.1.6	Posterior Tibial Pulses	14-105				
14.2.3.1.7	Leg Pulses	14-108				
14.2.3.1.8	Peripheral Pulses	14-110				
14.3 DISCUSSION	N	14-113				
14.4 SUMMARY		14-115				
14.4.1 Model	1: Group Analysis	14-115				
	2: Initial Dioxin Analysis					
14.4.3 Model	3: Categorized Dioxin Analysis	14-119				
14.4.4 Model	4: 1987 Dioxin Level Analysis	14-121				
14.5 CONCLUSIO	ON					
EFERENCES						

List of Tables

Table 14-1. Statistical Analysis for the Cardiovascular Assessment	14-8
Table 14-2. Number of Participants Excluded or with Missing Data for the Cardiovascular	1411
Assessment	
Table 14-3. Analysis of Essential Hypertension	
Table 14-4. Analysis of Heart Disease (Excluding Essential Hypertension)	
Table 14-5. Analysis of Myocardial Infarction	
Table 14-6. Analysis of Stroke or Transient Ischemic Attack	
Table 14-7. Analysis of Systolic Blood Pressure (Continuous)	
Table 14-8. Analysis of Systolic Blood Pressure (Discrete)	
Table 14-9. Analysis of Diastolic Blood Pressure (Continuous)	14-35
Table 14-10. Analysis of Diastolic Blood Pressure (Discrete)	14-38
Table 14-11. Analysis of Heart Sounds	14-40
Table 14-12. Analysis of Overall Electrocardiograph (ECG)	14-43
Table 14-13. Analysis of Right Bundle Branch Block	14-45
Table 14-14. Analysis of Left Bundle Branch Block	14-47
Table 14-15. Analysis of Non-Specific ST- and T-Wave Changes	14-50
Table 14-16. Analysis of Bradycardia	14-52
Table 14-17. Analysis of Tachycardia	14-55
Table 14-18. Analysis of Arrhythmia	14-58
Table 14-19. Analysis of Evidence of Prior Myocardial Infarction	14-60
Table 14-20. Analysis of ECG: Other Diagnoses	14-63
Table 14-21. Analysis of Funduscopic Examination	14-66
Table 14-22. Analysis of Carotid Bruits	14-69
Table 14-23. Analysis of Radial Pulses	14-71
Table 14-24. Analysis of Femoral Pulses	14-74
Table 14-25. Analysis of Popliteal Pulses	14-76
Table 14-26. Analysis of Dorsalis Pedis Pulses	14-79
Table 14-27. Analysis of Posterior Tibial Pulses	14-81
Table 14-28. Analysis of Leg Pulses	14-83
Table 14-29. Analysis of Peripheral Pulses	14-86
Table 14-30. Analysis of ICVI Index	14-88
Table 14-31. Longitudinal Analysis of Systolic Blood Pressure (mm Hg) (Continuous)	14-92
Table 14-32. Longitudinal Analysis of Systolic Blood Pressure (Discrete)	14-95
Table 14-33 Longitudinal Analysis of Femoral Pulses	14-98

Table 14-34.	Longitudinal Analysis of Popliteal Pulses	.14-101
Table 14-35.	Longitudinal Analysis of Dorsalis Pedis Pulses	.14-103
Table 14-36.	Longitudinal Analysis of Posterior Tibial Pulses	.14-106
Table 14-37.	Longitudinal Analysis of Leg Pulses	.14-108
Table 14-38.	Longitudinal Analysis of Peripheral Pulses	.14-111
Table 14-39.	Summary of Group Analysis (Model 1) for Cardiovascular Variables (Ranch Hands vs. Comparisons)	.14-115
Table 14-40.	Summary of Initial Dioxin Analysis (Model 2) for Cardiovascular Variables (Ranch Hands Only)	.14-118
Table 14-41.	Summary of Categorized Dioxin Analysis (Model 3) for Cardiovascular Variables (Ranch Hands vs. Comparisons)	.14-119
Table 14-42.	Summary of 1987 Dioxin Analysis (Model 4) for Cardiovascular Variables (Ranch Hands Only)	.14-121

14 CARDIOVASCULAR ASSESSMENT

14.1 INTRODUCTION

14.1.1 Background

Animal research into the cardiotoxicity of 2,3,7,8-tetrachlorodibenzo-p-dioxin (dioxin) has focused on acute biochemical and functional abnormalities associated with high-level exposure. In one study (1), rats were found to have reductions in pulse and blood pressure 6 days after administration of $40 \mu g/kg$ of dioxin by gavage and were less responsive to the chronotropic effect of isoproterenol, a beta-agonist. The authors of the study, noting a 66-percent reduction in serum thyroxine, postulated a down regulation of beta-receptors associated with the hypothyroid state rather than a direct cardiotoxic effect. Their findings were consistent with other studies that documented changes in myocardial beta-receptors with reduced serum indices of thyroid function and decreased beta-adrenergic responsiveness to isoproterenol in the ventricular papillary muscle of guinea pigs (2). Experiments into the effects of dioxin on myocardial contractility in rat (3) and guinea pig (4) atrial muscle have yielded mixed results; the primary cardiotoxic effects remain uncertain.

The biochemical effects of dioxin on cardiac muscle have been the subject of several reports. An increase in lipid peroxidation and a decrease in superoxide dismutase activity were noted in the hearts of female rats after dioxin administration (1). Dose-dependent decreases in adipose tissue lipoprotein lipase activity and hepatic low-density lipoprotein binding occurred in rabbits (5) and other laboratory animals (6) in association with elevated serum triglycerides. Electron microscopic studies have documented pre-atherosclerotic lesions in the aortic arch in association with these biochemical abnormalities (5) and dioxin exposure has been associated with intravascular thrombosis in rats (7). Two recent studies provide evidence that the developing vascular endothelium of fish embryos may be a target organ for dioxin toxicity (8, 9).

Numerous studies have focused on the effects of dioxin toxicity on lipid metabolism in experimental animals and may be relevant to herbicide exposure as a risk factor for the development of heart disease in man. Dioxin-induced hyperlipidemia has been documented in rats (10, 11), guinea pigs (12), and rabbits (5).

Numerous epidemiological studies have investigated cardiovascular mortality and morbidity in populations exposed to dioxin by occupation and consequent to industrial accidents (13–22). Other reports have examined similar endpoints in veterans who served in the Vietnam War (23–35). Some occupational (13, 20) and veterans' studies (23, 25, 26, 28–31) cited have shown no increase in cardiovascular mortality associated with exposure to dioxin, and several have documented a significant reduction in risk (23, 26, 27). However, in the 1994 Air Force Health Study (AFHS) mortality update (36), the Ranch Hand nonflying enlisted personnel were found to be at higher risk for death associated with circulatory disease than the Comparison nonflying enlisted personnel. Most occupational studies have found no increased risk for the development of cardiovascular disease related to dioxin exposure (13–16, 20). In two reports of the 1976 Seveso, Italy, industrial accident, dioxin exposure was associated with statistically significant increases in mortality because of coronary, cerebrovascular, and hypertensive vascular disease (18, 19).

The latest morbidity follow-up study of BASF Corporation employees highly exposed to dioxin during a chemical reactor incident in 1953 has been published (21). Almost half of the study group had

extrapolated serum dioxin levels of more than 1,000 parts per trillion (ppt). Across all exposure categories, there was no significant increase in the incidence of ischemic heart disease.

A more recently published retrospective cohort study examined cardiovascular mortality in 1,189 German chemical workers who had significant dioxin exposure in the 1950s (37). In this study, exposure was verified and subjects stratified into deciles based on serum and adipose tissue dioxin levels. There was a slight reduction in mortality risk at the two lowest levels of exposure, but a clear pattern of increasing risk for all-cause cardiovascular mortality and, particularly, for that associated with ischemic heart disease. The dose-response trend for both causes of mortality was significant (p≤0.01).

The well-established roles of diabetes mellitus and lipid disorders as risk factors in the development of cardiovascular disease have generated considerable interest in the potential intermediary role these metabolic indices might have on cardiovascular outcomes associated with dioxin exposure. Data and results from this (35, 38) and other epidemiological studies (22, 37, 39–44) are considered in the Gastrointestinal Assessment chapter (Chapter 13) and the Endocrine Assessment chapter (Chapter 16).

Previous AFHS examinations have shown mixed results with respect to cardiovascular endpoints. In the baseline and 1987 follow-up examinations, manual examination of the pulses revealed an increased prevalence of pulse deficits in the Ranch Hand cohort relative to Comparisons (45, 46), results noted as well in studies of residents exposed to dioxin in Times Beach, Missouri (47, 48). In the 1985 AFHS follow-up examination, which incorporated Doppler peripheral vascular studies into the protocol, no significant group differences were found (49). When the 1987 examination data were analyzed relative to serum dioxin levels, Ranch Hand participants in one high exposure category had higher percentages of peripheral pulse abnormalities by manual examination than did Comparisons (34). In addition, Ranch Hands with the highest current dioxin levels were at greater risk for the development of systemic arterial hypertension than were Comparisons. In contrast, there was a significant reduction in risk for the development of heart disease reported historically or by a verified medical records review.

In the 1992 follow-up examination, Ranch Hands were more likely than Comparisons to have elevated systolic blood pressures, and through 1990, there was an increase in cardiovascular disease mortality in the nonflying enlisted personnel. However, surviving Ranch Hands overall were found to be less at risk for the development of heart disease over time, and a significant inverse dose-response effect was noted with respect to the current body burden of dioxin (35).

14.1.2 Summary of Previous Analyses of the Air Force Health Study

14.1.2.1 1982 Baseline Study Summary Results

The 1982 baseline examination found no statistically significant differences between the Ranch Hand and Comparison groups in systolic or diastolic blood pressure, the frequency of abnormal electrocardiographs (ECGs), heart sound abnormalities, abnormal funduscopic findings, or carotid bruits. A statistically significant difference emerged in the frequency of abnormal peripheral pulses: 12.8 percent of the non-Black Ranch Hands exhibited absent or diminished peripheral pulses, compared to 9.4 percent of the non-Black Original Comparisons (p=0.05). No statistically significant differences were found between the two groups in the occurrence of reported or verified heart disease or heart attacks.

Greater than 80 percent of the cardiac conditions reported on the study questionnaire were verified by a detailed review of medical records. There was also a strong correlation between the past medical history of cardiac disease and the baseline examination cardiovascular findings, although the differences in peripheral pulse abnormalities occurred primarily in older individuals without a history of cardiovascular

disease. Finally, the well-known risk factors of age, smoking, and cholesterol were found to be correlated with each other and with several of the cardiovascular response variables.

14.1.2.2 1985 Follow-up Study Summary Results

The analysis of cardiovascular disease history did not reveal significant group differences in reported or verified hypertension, reported heart disease, or reported or verified heart attacks. There were no group differences in verified heart disease. The verified cardiovascular history and the central and peripheral cardiovascular abnormalities detected at the physical examination were correlated, supporting accuracy and validity of the cardiovascular measurements.

In the analyses of peripheral vascular function, no significant overall group differences were observed for abnormalities involving radial, femoral, popliteal, posterior tibial, dorsalis pedis, or three anatomic aggregates of these pulses (leg pulses, peripheral pulses, and all pulses), either by manual palpation or Doppler techniques. This overall finding was in distinct contrast to the 1982 baseline examination, which, by the manual palpation method, showed significant peripheral pulse deficits in Ranch Hands. This reversal in pulse findings over the two examinations may be attributed to the rigid 4-hour tobacco abstinence applied prior to Doppler testing, although other factors may have been involved.

14.1.2.3 1987 Follow-up Study Summary Results

The assessment of the central cardiac function also found the groups to be similar, although significantly fewer Ranch Hands than Comparisons had bradycardia and more Ranch Hands than Comparisons had arrhythmias (marginally significant).

For the peripheral vascular function, Ranch Hands had a higher or marginally higher mean or percent abnormal for diastolic blood pressure (continuous form), carotid bruits, femoral pulses, and dorsalis pedis pulses than did Comparisons. No difference between the two groups was detected in the discrete analysis of diastolic blood pressure. The percentage of radial pulse abnormalities was marginally higher in Comparisons than in Ranch Hands. On the three pulse indices (leg, peripheral, and all pulses), Ranch Hands had marginally or significantly higher percentages of abnormalities than did Comparisons.

14.1.2.4 Serum Dioxin Analysis of 1987 Follow-up Study Summary Results

The cardiovascular evaluation found a marginally significant association between initial dioxin and a decrease in the reported history of heart disease, and a significant negative association with verified history of heart disease. In addition, the analyses of categorized current dioxin also indicated a decrease in verified history of heart disease for Ranch Hands with the highest current dioxin levels relative to Comparisons with background levels. These Ranch Hands also had more essential hypertension by history (after removing the variables body fat and cholesterol from the model).

The analyses of the peripheral vascular function variables displayed significantly higher mean levels of diastolic blood pressure for Ranch Hands in the low and high categories than Comparisons (without adjustment for body fat). Similar to the analysis of systolic blood pressure, the discretized analysis of diastolic blood pressure did not display a significant association with dioxin within the low and high current dioxin categories. Ranch Hands generally exhibited a significant or marginally significant higher risk of absent femoral, dorsalis pedis, and posterior tibial pulses relative to Comparisons. These observations could represent a subclinical effect and emphasize the importance of continued follow-up and evaluation.

14.1.2.5 1992 Follow-up Study Summary Results

The cardiovascular evaluation found a marginally significant group difference for verified heart disease, excluding essential hypertension for enlisted flyers with Ranch Hands having a greater history of heart disease than Comparisons. Similar to the 1987 study, verified heart disease decreased significantly for increasing levels of current dioxin. Ranch Hands also displayed an increased history of essential hypertension for increasing levels of current dioxin.

A few other central cardiac function endpoints, including non-specific ST- and T-wave changes, right bundle branch block, and prior ECG evidence of myocardial infarction, displayed significant positive associations with current dioxin; none of these endpoints also displayed any group difference between Ranch Hands and Comparisons. These findings, in conjunction with the increase in the number of deaths caused by diseases of the circulatory system for Ranch Hand nonflying enlisted personnel based on the 1994 AFHS mortality update (34), showed potential associations with dioxin requiring further observation.

The analyses of the peripheral vascular function variables displayed significant group differences for the enlisted groundcrew stratum for a few of the pulse endpoints and significant differences between Ranch Hands in the high dioxin category and Comparisons. None of these associations was reinforced by a significant association with initial or current dioxin. Longitudinal analyses of the pulse endpoints also indicated that Ranch Hands in the enlisted groundcrew stratum and in the high initial dioxin category had a greater prevalence of pulse deficits since the 1985 follow-up examination than Comparisons. Again, these associations were not reinforced by a significant dose-response effect with initial dioxin.

In general, after reviewing the results of the cardiovascular assessment as a whole, the development of cardiovascular disease did not appear to be associated positively with dioxin. Dioxin associations with selected endpoints, as discussed above, together with mortality results, pointed to the need for further evaluation.

14.1.3 Parameters for the 1997 Cardiovascular Assessment

14.1.3.1 Dependent Variables

The analysis of the cardiovascular assessment was based on data collected from the 1997 questionnaire and physical examination and subsequent medical records verification. No laboratory examination data were analyzed as cardiovascular dependent variables, although data from the laboratory examination were used as covariates.

14.1.3.1.1 Medical Records Data

During the baseline, 1985, 1987, and 1992 AFHS examination health interviews, each participant was asked whether he had a heart condition. Medical records were sought to verify all reported conditions and to determine the time of occurrence of major cardiac events. In addition, the self-reported review-of-systems recorded the overall history of heart trouble and other serious illnesses. Data collected in a similar fashion at the 1997 follow-up was verified and combined with data from the four previous examinations to create a lifetime history for four conditions: essential hypertension, heart disease (excluding essential hypertension), myocardial infarction, and stroke or transient ischemic attack. Each of these conditions was classified as "yes" or "no" and analyzed.

International Classification of Diseases, 9th Revision, Clinical Modification (ICD-9-CM) codes were used to construct the four conditions described above. The following ICD-9-CM codes were used: essential hypertension (ICD-9-CM codes 401.0-401.9), heart disease (excluding essential hypertension)

(ICD-9-CM codes 391.0-391.9, 392.0, 393.0-398.99, 402.0-402.91, 404.0-404.9, 410.0-417.9, and 420.0-429.9), myocardial infarction (ICD-9-CM codes 410.0-410.9, and 412), and stroke or transient ischemic attack (ICD-9-CM codes 435.0-436).

Participants with a verified pre-SEA heart condition were excluded from all analyses. A pre-SEA heart condition included pre-SEA myocardial infarction, but did not include pre-SEA essential hypertension. Participants with a verified pre-SEA history of essential hypertension also were excluded from the analysis of verified history of essential hypertension.

14.1.3.1.2 Physical Examination Data and Self-reported Questionnaire Data

Cardiovascular data analyzed from the 1997 physical examination were divided into two main categories: central cardiac function and peripheral vascular function.

14.1.3.1.2.1 Central Cardiac Function

The assessment of the central cardiac function at the cardiovascular examination was made by measurements of systolic blood pressure, diastolic blood pressure, heart sounds (by auscultation), and an ECG. Systolic and diastolic blood pressure were determined by a Critikon Dinamap 1846SXP® automated electronic monitor with the nondominant arm placed at heart level; the lowest diastolic pressure and the corresponding systolic pressure were recorded. Detection of abnormal heart sounds was conducted by standard auscultation with the participant placed in sitting, supine, and left lateral supine positions. Fourth heart sounds were assessed: murmurs were graded in intensity and location and were judged by the examiners to be functional (normal) or organic (abnormal) in nature. The standard 12-lead ECG was performed, and an additional strip in limb lead II was produced if any arrhythmia was found. Participants were asked to abstain from tobacco for at least 4 hours prior to the ECG because of the arterial constrictive effect of nicotine. The following items were considered to be abnormal: right bundle branch block, left bundle branch block, nonspecific ST- and T-wave changes, bradycardia (a resting pulse rate less than 50 beats per minute), tachycardia (a resting pulse rate greater than 100 beats per minute), arrhythmia (any irregularity of heart rhythm including premature beats but excluding normal sinus rhythm), evidence of a prior myocardial infarction, and other diagnoses (e.g., ventricular aneurysm, Wolff-Parkinson-White syndrome). Some arrhythmias (e.g., atrial flutter, atrial fibrillation, and junctional rhythm) required more evaluation and surveillance than others, but all were grouped together for evaluation in this study.

Variables analyzed in the evaluation of the central cardiac function included systolic blood pressure, diastolic blood pressure, heart sounds, an overall ECG assessment, and eight conditions associated with the ECG. These eight conditions were right bundle branch block, left bundle branch block, nonspecific ST- and T-wave changes, bradycardia, tachycardia, arrhythmia, evidence of a prior myocardial infarction, and other diagnoses. Both systolic and diastolic blood pressure were analyzed as a continuous variable and also as a discrete variable. Systolic blood pressure was classified as "normal" (≤140 mm Hg) and "high" (>140 mm Hg), and diastolic blood pressure was classified as "normal" (≤90 mm Hg) and "high" (>90 mm Hg). Participants with a verified pre-SEA heart condition were excluded from all analyses of the central cardiac function variables.

14.1.3.1.2.2 Peripheral Vascular Function

The peripheral vascular function was assessed during the cardiovascular examination by funduscopic examination of small vessels; presence or absence of carotid bruits; determination of the radial, femoral, popliteal, dorsalis pedis, and posterior tibial pulses by Doppler techniques; and a measure of intermittent claudication and vascular insufficiency.

The funduscopic examination was conducted with undilated pupils in a standard manner, with emphasis placed upon the detection of increased light reflex, arteriovenous nicking (a sign of chronic blood pressure elevation), hemorrhages, exudates, papilledema, and arteriolar spasm. The presence or absence of carotid bruits was assessed by auscultation over both carotid arteries.

The Doppler procedure for examining pulses is a progressive array of measurements designed to determine whether a pulse abnormality exists, where the obstruction is most likely located, and whether it has functional implications. The determination of a pulse abnormality was based upon an analysis of recorded Doppler waveform morphology. Pulsatility, systolic forward flow, diastolic reverse flow, and diastolic oscillations were examined.

The funduscopic examination, carotid bruits, and the five pulses also were dichotomized as "abnormal" or "normal" (or "presence" or "absence") and analyzed. Pulses were considered abnormal if no arterial flow or a monophasic arterial flow was present on either side. In addition, two pulse indices were constructed from the radial, femoral, popliteal, dorsalis pedis, and posterior tibial pulse measurements as follows:

- Leg pulses: femoral, popliteal, dorsalis pedis, and posterior tibial pulses
- Peripheral pulses: radial, femoral, popliteal, dorsalis pedis, and posterior tibial pulses.

Each of these indices was considered "normal" if all components were normal and "abnormal" if one or more pulses were abnormal.

In the 1997 questionnaire, each participant was asked the following questions:

- Do you get a pain in either or both of your legs while walking?
- Does this pain ever begin when you are standing still or sitting?
- Do you get this pain in either or both of your calf muscles?

The self-reported answers were used to detect intermittent claudication and vascular insufficiency (yes, no), which indicate an insufficient oxygen supply to the leg muscles. A participant was judged to have intermittent claudication and vascular insufficiency if he answered "yes" to the first and third questions and "no" to the second question. Participants with a verified pre-SEA heart condition were excluded from all analyses of the peripheral vascular function variables.

14.1.3.2 Covariates

A number of covariates were examined for inclusion in the adjusted analysis of the cardiovascular assessment. Many of these covariates are considered to be classical risk factors for chronic heart disease. Covariates examined included age, race, military occupation, lifetime alcohol history, current alcohol use, lifetime cigarette smoking history, current level of cigarette smoking, cholesterol, high-density lipoprotein (HDL), cholesterol-HDL ratio, body fat, personality type, family history of heart disease, family history of heart disease before the age of 45, diabetic class, and current use of blood pressure medication (for the blood pressure variables).

Age, race, and military occupation were determined from military records. Lifetime alcohol history was based on information from the 1997 questionnaire and combined with similar information gathered at the 1987 and 1992 follow-up examinations. Each participant was asked about his drinking patterns throughout his lifetime. When a participant's drinking patterns changed, he was asked to describe how his alcohol consumption differed and the duration of time that the drinking pattern lasted. The participant's average daily alcohol consumption was determined for each of the reported drinking pattern

periods throughout his lifetime, and an estimate of the corresponding total number of drink-years was derived. One drink-year was the equivalent of drinking 1.5 ounces of an 80-proof alcoholic beverage, one 12-ounce beer, or one 5-ounce glass of wine per day for 1 year.

Current cigarette smoking and lifetime cigarette smoking history were based on questionnaire data. For lifetime cigarette smoking history, the respondent's average smoking was estimated over his lifetime based on his responses to the 1997 questionnaire, with 1 pack-year defined as 365 packs of cigarettes smoked during a single year.

Cholesterol, HDL, and the cholesterol-HDL ratio were based on 1997 laboratory measurements. Body fat was calculated from a metric body mass index (50); the formula is

Body Fat (in percent) =
$$\frac{Weight(kg)}{[Height(m)]^2} \bullet 1.264 - 13.305.$$

Personality type was determined from the Jenkins Activity Survey administered during the 1997 follow-up examination and was derived from a discriminant-function equation based on questions that best discriminate men judged to be type A from those judged to be type B (51). Positive scores reflected the type A direction and negative scores reflected the type B direction. Personality type was dichotomized as type A or type B.

Family history of heart disease was defined as "yes" if the participant's mother, father, sister(s), or brother(s) had heart trouble or heart disease and "no" otherwise. Family history of heart disease before the age of 45 was defined as "yes" if the participant's mother, father, sister(s), or brother(s) had heart trouble or heart disease before the age of 45 and "no" otherwise. Blood pressure medication (yes, no) was used as a covariate for the adjusted analysis of the systolic and diastolic blood pressure variables only.

Diabetic class was used as a covariate in the analysis of the 1997 follow-up. Diabetes is a known risk factor for cardiovascular disease. In the 1997 questionnaire, a general screening question on diabetes was posed. Each participant was asked during the in-person health interview the following question: "Since the date of the last interview, has a doctor told you for the first time that you had diabetes?" All affirmative responses were verified by a medical records review and added to previously reported and verified information on diabetes from the 1982 baseline and the 1985, 1987, and 1992 follow-up examinations for each participant. Participants with a verified history of diabetes were combined with those participants with a 2-hour postprandial glucose level of 200 mg/dl or greater at the 1997 physical examination and classified as "diabetic" for the diabetic class covariate. Those participants without a verified history of diabetes and with a 2-hour postprandial glucose level of less than 200 mg/dl at the 1997 physical examination were classified as either "impaired" (140 mg/dl ≤ 2-hour postprandial glucose < 200 mg/dl) or "normal" (2-hour postprandial glucose < 140 mg/dl).

The current use of blood pressure medication was used as a covariate for the adjusted analysis of systolic and diastolic blood pressures. This information was reported by the participant on a self-reported form that listed physicians and medications, and through a question in the in-person interview.

The following dependent variables—essential hypertension, heart disease excluding essential hypertension, myocardial infarction, and stroke or transient ischemic attack—capture a history of a cardiovascular condition rather than the current state of a participant's life at the time of the physical examination. Consequently, to reflect the historical nature of these dependent variables, lifetime alcohol history and lifetime cigarette smoking history were used as covariates, but current alcohol use and current cigarette smoking were not. Lifetime alcohol history and lifetime cigarette smoking history reflect the

cumulative lifetime effects of alcohol use and tobacco, respectively, whereas current alcohol use and current cigarette smoking emphasize the short period of time near the date of the physical examination.

14.1.4 Statistical Methods

Table 14-1 summarizes the statistical analysis performed for the cardiovascular assessment. The first part of this table describes the dependent variables and identifies the covariates and the statistical methods. The second part of this table further describes the covariates. A covariate was used in its continuous form whenever possible for all adjusted analyses. If a covariate was inherently discrete (e.g., military occupation), or if a categorized form was needed to develop measures of association with the dependent variables, the covariate was categorized as shown in Table 14-1.

Table 14-2 provides a summary of the number of participants with missing dependent variable or covariate data. In addition, the number of participants excluded from analysis is given.

Table 14-1. Statistical Analysis for the Cardiovascular Assessment

Dependent Variables

Variable (Units)	Data Source	Data Form	Cutpoints	Covariates ^a	Exclusions ^b	Statistical Analysis and Methods
Essential Hypertension	MR-V	D	Yes No	(1)	(a)	U:LR A:LR
Heart Disease (Excluding Essential Hypertension)	MR-V	D	Yes No	(1)	(b)	U:LR A:LR
Myocardial Infarction	MR-V	D	Yes No	(1)	(b)	U:LR A:LR
Stroke or Transient Ischemic Attack	MR-V	D	Yes No	(1)	(b)	U:LR,CS A:LR
Systolic Blood Pressure (mm Hg)	PE	D/C	High: >140 Normal: ≤140	(2)	(b)	U:LR,GLM A:LR,GLM L:LR,GLM
Diastolic Blood Pressure (mm Hg)	PE	D/C	High: >90 Normal: ≤90	(2)	(b)	U:LR,GLM A:LR,GLM
Heart Sounds	PE	D	Abnormal Normal	(3)	(b)	U:LR A:LR
Overall Electrocardiograph (ECG)	PE	D	Abnormal Normal	(3)	(b)	U:LR A:LR
ECG: Right Bundle Branch Block	PE	D	Yes No	(3)	(b)	U:LR A:LR
ECG: Left Bundle Branch Block	PE	D	Yes No	(3)	(b)	U:LR,CS A:LR
ECG: Non-specific ST-and T-Wave Changes	PE	D	Yes No	(3)	(b)	U:LR A:LR
ECG: Bradycardia	PE	D	Yes No	(3)	(b)	U:LR A:LR
ECG: Tachycardia	PE	D	Yes No	(3)	(b)	U:LR,CS A:LR

Table 14-1. Statistical Analysis for the Cardiovascular Assessment (Continued)

Variable (Units)	Data Source	Data Form	Cutpoints	Covariates ^a	Exclusions ^b	Statistical Analysis and Methods
ECG: Arrhythmia	PE	D	Yes No	(3)	(b)	U:LR A:LR
ECG: Evidence of Prior Myocardial Infarction	PE	D	Yes No	(3)	(b)	U:LR A:LR
ECG: Other Diagnoses	PE	D	Yes No	(3)	(b)	U:LR,CS A:LR
Funduscopic Examination	PE	D	Abnormal Normal	(3)	(b)	U:LR A:LR
Carotid Bruits	PE	D	Present Absent	(3)	(b)	U:LR A:LR
Radial Pulses	PE	D	Abnormal Normal	(3)	(b)	U:LR A:LR
Femoral Pulses	PE	D	Abnormal Normal	(3)	(b)	U:LR A:LR L:LR
Popliteal Pulses	PE	D	Abnormal Normal	(3)	(b)	U:LR A:LR L:LR
Dorsalis Pedis Pulses	PE	D	Abnormal Normal	(3)	(b)	U:LR A:LR L:LR
Posterior Tibial Pulses	PE	D	Abnormal Normal	(3)	(b)	U:LR A:LR L:LR
Leg Pulses	PE	D	Abnormal Normal	(3)	(b)	U:LR A:LR L:LR
Peripheral Pulses	PE	D	Abnormal Normal	(3)	(b)	U:LR A:LR L:LR
Intermittent Claudication and Vascular Insufficiency (ICVI) Index	Q-SR	D	Abnormal Normal	(3)	(b)	U:LR A:LR

^aCovariates:

^bExclusions:

^{(1):} age, race, military occupation, lifetime cigarette smoking history, lifetime alcohol history, cholesterol, HDL, cholesterol-HDL ratio, diabetic class, body fat, personality type, family history of heart disease, family history of heart disease before age 45.

^{(2):} age, race, military occupation, lifetime cigarette smoking history, current cigarette smoking, lifetime alcohol history, current alcohol use, cholesterol, HDL, cholesterol-HDL ratio, diabetic class, body fat, personality type, family history of heart disease, family history of heart disease before age 45, taking blood pressure medication. (3): age, race, military occupation, lifetime cigarette smoking history, current cigarette smoking, lifetime alcohol history, current alcohol use, cholesterol, HDL, cholesterol-HDL ratio, diabetic class, body fat, personality type, family history of heart disease, family history of heart disease before age 45.

⁽a): participants with a pre-SEA heart condition, participants with pre-SEA essential hypertension.

⁽b): participants with a pre-SEA heart condition.

Table 14-1. Statistical Analysis for the Cardiovascular Assessment (Continued)

Covariates

Variable (Units)	Data Source	Data Form	Cutpoints
Age (years)	MIL	D/C	Born ≥1942 Born <1942
Race	MIL	D	Black Non-Black
Occupation	MIL	D	Officer Enlisted Flyer Enlisted Groundcrew
Lifetime Alcohol History (drink-years)	Q-SR	D/C	0 >0–40 >40
Current Alcohol Use (drinks/day)	Q-SR	D/C	0-1 >1-4 >4
Lifetime Cigarette Smoking History (pack-years)	Q-SR	D/C	0 >0–10 >10
Current Cigarette Smoking (cigarettes/day)	Q-SR	D/C	0-Never 0-Former >0-20 >20
Cholesterol (mg/dl)	LAB	D/C	≤200 >200–239 >239
High Density Lipoprotein (mg/dl)	LAB	D/C	0–35 >35
Cholesterol-HDL Ratio	LAB	D/C	0–5 >5
Body Fat (percent)	PE	D/C	Obese: >25% Lean or Normal: ≤25%
Personality Type	PE	D	A direction B direction
Family History of Heart Disease	Q-SR	D	Yes No
Family History of Heart Disease Before Age 45	Q-SR	D	Yes No
Diabetic Class	LAB/MR-V	D	 Diabetic: past history or ≥200 mg/dl 2-hr. postprandial glucose Impaired: 140–<200 mg/dl 2-hr. postprandial glucose Normal: <140 mg/dl 2-hr. postprandial glucose
Taking Blood Pressure Medication	Q-SR/MR-V	D	Yes No

Table 14-1. Statistical Analysis for the Cardiovascular Assessment (Continued)

Abbreviations

Data Source: LAB: 1997 laboratory results

MIL: Air Force military records MR-V: Medical records (verified) PE: 1997 physical examination

Q-SR: Health questionnaires (self-reported)

Data Form: D: Discrete analysis only

D/C: Discrete and continuous analyses for dependent variables; appropriate form for analysis

(either discrete or continuous) for covariates

Statistical Analysis: U: Unadjusted analysis

A: Adjusted analysisL: Longitudinal analysis

Statistical Methods: CS: Chi-square contingency table analysis (continuity-adjusted)

GLM: General linear models analysis LR: Logistic regression analysis

Table 14-2. Number of Participants Excluded or with Missing Data for the Cardiovascular Assessment

		Group	(Ranch Ha	nds Only)	Catego	orized Dioxin
Variable	Ranch				Ranch	
Use	Hand	Comparison	Initial	1987	Hand	Comparison
DEP	1	1	0	1	1	1
DEP	0	1	0	0	0	1
DEP	0	2	0	0	0	2
DEP	0	2	0	0	0	2
DEP	0	4	0	0	0	4
DEP	0	4	0	0	0	4
DEP	0	4	0	0	0	4
DEP	1	0	0	1	1	0
COV	6	2	3	6	6	1
COV	1	0	0	1	1	0
COV	2	1	1	2	2	1
COV	1	0	0	1	1	0
COV	1	1	1	1	1	1
COV	1	1	1	1	1	1
COV	3	0	1	3	3	0
COV	10	6	5	10	10	6
COV	22	22	11	22	22	21
EXC	9	18	5	7	7	17
	Use DEP DEP DEP DEP DEP DEP DEP DEP COV COV COV COV COV COV COV COV	Variable Use Ranch Hand DEP 1 DEP 0 DEP 0 DEP 0 DEP 0 DEP 0 DEP 0 DEP 1 COV 6 COV 1 COV 2 COV 1 COV 1 COV 3 COV 10 COV 22	Use Hand Comparison DEP 1 1 DEP 0 1 DEP 0 2 DEP 0 2 DEP 0 4 DEP 0 4 DEP 0 4 DEP 1 0 COV 6 2 COV 1 0 COV 2 1 COV 1 1 COV 1 1 COV 1 1 COV 3 0 COV 10 6	Variable Use Ranch Hand Hand Comparison Initial DEP 1 1 0 DEP 0 1 0 DEP 0 2 0 DEP 0 2 0 DEP 0 4 0 DEP 0 4 0 DEP 0 4 0 DEP 0 4 0 DEP 1 0 0 COV 6 2 3 COV 1 0 0 COV 2 1 1 COV 1 1 1 COV 1 1 1 COV 3 0 1 COV 10 6 5	Variable Use Ranch Hand Comparison Initial 1987 DEP 1 1 0 1 DEP 0 1 0 0 DEP 0 2 0 0 DEP 0 2 0 0 DEP 0 4 0 0 DEP 1 0 0 1 COV 6 2 3 6 COV 1 0 0 1 COV 2 1 1 2 COV 1 0 0 1 COV 1 1 1 1 COV 1 1 1 1 COV 3 0 1 3 COV	Variable Use Ranch Hand Comparison Initial 1987 Ranch Hand DEP 1 1 0 1 1 DEP 0 1 0 0 0 DEP 0 2 0 0 0 DEP 0 2 0 0 0 DEP 0 4 0 0 0 DEP 1 0 1 1 COV 6 2 3 6 6 COV 1 0 0 1 1 COV 1 0 0 1 1 COV 1 1 1 1 1 COV 1 1 1 1 1

Table 14-2. Number of Participants Excluded or with Missing Data for the Cardiovascular Assessment (Continued)

		Dioxin Group (Ranch Hands Only) Categorized Dioxin			orized Dioxin		
Variable	Variable Use	Ranch Hand	Comparison	Initial	1987	Ranch Hand	Comparison
Pre-SEA Heart Condition	EXC	11	19	6	11	11	18
Pre-SEA Essential Hypertension	EXC	11	14	7	11	11	14

Note: DEP = Dependent variable.

COV = Covariate. EXC = Exclusion.

870 Ranch Hands and 1,251 Comparisons.

482 Ranch Hands for initial dioxin; 863 Ranch Hands for 1987 dioxin. 863 Ranch Hands and 1,213 Comparisons for categorized dioxin.

14.1.4.1 Longitudinal Analysis

The cardiovascular longitudinal analysis was based on the association of exposure with changes in systolic blood pressure between the 1982 and 1997 examinations and six pulse measurements between the 1985 and 1997 examinations. The longitudinal analysis for systolic blood pressure was based on this variable in both the continuous and discrete forms. The six pulse measurements included femoral pulses, popliteal pulses, dorsalis pedis pulses, posterior tibial pulses, leg pulses, and peripheral pulses. The 1985 and 1997 measurements were used for the pulse assessments because the Doppler assessment of pulses was conducted at these two examinations and was not conducted at the 1982 baseline examination.

14.2 RESULTS

14.2.1 Dependent Variable-Covariate Associations

The associations between the dependent variables examined in the cardiovascular assessment and the covariates used in the adjusted analysis were investigated; the results are presented in Appendix F, Table F-6. These associations are pairwise between the dependent variable and the covariate and are not adjusted for any other covariates. Participants with a pre-SEA heart condition were excluded from all analyses. In addition, participants with pre-SEA essential hypertension were excluded from the analysis of essential hypertension.

Tests of covariate association showed age (p=0.001), lifetime alcohol history (p=0.001), cholesterol-HDL ratio (p=0.005), body fat (p=0.001), personality type (p=0.039), family history of heart disease (p=0.001), family history of heart disease before age 45 (p=0.003), and diabetic class (p=0.001) to be significantly associated with essential hypertension. Older participants had more essential hypertension than did younger participants (48.0% versus 32.9%). Essential hypertension was highest for the heaviest drinkers (in terms of drink-years) (48.2%), followed by participants who did not drink (39.0%), then moderate drinkers (38.5%). Essential hypertension increased with the cholesterol-HDL ratio and body fat. Participants with personality type B had a higher percentage of essential hypertension than did type A participants (43.0% versus 38.4%). Essential hypertension occurred more often for participants who had a family history of heart disease and for participants who had a family history of heart disease before age 45. Essential hypertension was greatest for diabetics (59.4%), followed by participants in the impaired diabetic class (52.4%), then participants classified as normal (34.6%).

Heart disease (excluding essential hypertension) was significantly associated with age (p=0.001), occupation (p=0.001), cholesterol (p=0.001), family history of heart disease (p=0.001), family history of heart disease before age 45 (p=0.018), and diabetic class (p=0.009). Heart disease increased with age and decreased with cholesterol level. Officers had the highest percentage of heart disease (68.7%), followed by enlisted flyers (66.6%), then enlisted groundcrew (56.7%). Participants with a family history of heart disease had more heart disease (66.6% versus 57.4%). Likewise, participants with a family history of heart disease before age 45 had more heart disease (69.9% versus 62.0%). Diabetic participants had the most heart disease (69.5%), followed by participants in the impaired diabetic class (64.1%), then participants classified as normal (60.8%).

The percentage of participants with a history of a myocardial infarction increased significantly with age (p=0.001) and lifetime cigarette smoking history (p=0.001), while decreasing significantly with cholesterol (p=0.001) and HDL cholesterol (p=0.012). The association with diabetic class was also significant (p=0.001). Participants in the normal diabetic class had the lowest percentage of participants with a myocardial infarction (6.8%), followed by participants in the impaired diabetic class (9.9%), then diabetics (14.2%).

Systolic blood pressure in its continuous form increased with age (p<0.001), lifetime alcohol history (p<0.001), lifetime cigarette smoking history (p=0.045), cholesterol (p=0.012), the cholesterol-HDL ratio (p=0.005), and body fat (p<0.001). Systolic blood pressure decreased significantly with current cigarette smoking (p=0.004). Tests of covariate associations also showed significant relations with occupation (p=0.005), diabetic class (p<0.001), and blood pressure medication (p<0.001). Enlisted flyers had the highest mean systolic blood pressure levels (127.1 mm Hg), followed by officers (126.1 mm Hg), then enlisted groundcrew (123.9 mm Hg). Participants in the normal diabetic class had the lowest mean systolic blood pressure levels (123.0 mm Hg), followed by participants in the impaired diabetic class (129.3 mm Hg), then diabetic participants (131.8 mm Hg). Participants taking blood pressure medication had a higher mean systolic blood pressure level (128.6 mm Hg) than those not taking blood pressure medication (123.9 mm Hg).

Systolic blood pressure in its dichotomous form increased with age (p=0.001), cholesterol (p=0.025), the cholesterol-HDL ratio (p=0.028), and body fat (p=0.001). Significant associations also were seen between systolic blood pressure and occupation (p=0.029), family history of heart disease (p=0.008), diabetic class (p=0.001), and blood pressure medication (p=0.001). Enlisted flyers had the greatest percentage of high systolic blood pressure values (23.6%), followed by officers (23.2%), then enlisted groundcrew (18.6%). Participants with a family history of heart disease had a greater prevalence of high systolic blood pressure values than did participants with no history of heart disease (23.3% versus 18.3%). Diabetic participants had the largest percentage of high systolic blood pressure values (31.9%), followed by participants in the impaired diabetic class (28.6%), then participants classified as normal (17.1%). Participants taking blood pressure medication had a greater prevalence of high systolic blood pressure values than participants not taking blood pressure medication (27.6% versus 18.5%).

Diastolic blood pressure in its continuous form decreased with age (p=0.009), lifetime cigarette smoking history (p=0.003), and current cigarette smoking (p=0.001). Diastolic blood pressure increased with cholesterol (p<0.001), the cholesterol-HDL ratio (p=0.004), and body fat (p<0.001). Race and diabetic class were also significantly associated with diastolic blood pressure (p=0.010 and p=0.030, respectively). Black participants had a higher mean diastolic blood pressure than non-Black participants (76.69 mm Hg versus 74.46 mm Hg). Participants in the impaired diabetic class had the highest mean diastolic blood pressure (75.94 mm Hg), followed by diabetic participants (74.41 mm Hg), then participants classified as normal (74.32 mm Hg).

Tests of covariate association for diastolic blood pressure in its discrete form showed significant relations with lifetime cigarette smoking history (p=0.003) and blood pressure medication (p=0.004). Moderate lifetime cigarette smokers (in terms of pack-years) had the greatest percentage of high diastolic blood pressure values (7.8%), followed by participants who never smoked and participants who were the heaviest smokers (4.1% each). Participants taking blood pressure medication had a greater prevalence of high diastolic blood pressure values than did participants not taking blood pressure medication (7.3% versus 4.1%).

The percentage of participants with abnormal heart sounds increased with age (p=0.001). Current cigarette smoking was also significantly associated with heart sounds (p=0.030). Former smokers had the highest prevalence of abnormal heart sounds (5.7%), followed by participants who smoked up to 20 cigarettes per day (3.4%), participants who smoked more than 20 cigarettes per day (2.9%), and participants who never smoked (2.9%).

The prevalence of abnormal overall ECG results increased with age (p=0.001) and body fat (p=0.008), while decreasing with cholesterol (p=0.041). Also significant were occupation (p=0.001), lifetime cigarette smoking history (p=0.002), current cigarette smoking (p=0.028), personality type (p=0.011), family history of heart disease (p=0.001), and diabetic class (p=0.001). Enlisted flyers had the highest percentage of abnormal overall ECG results (36.4%), followed by officers (34.6%), then enlisted groundcrew (26.3%). Heavy lifetime cigarette smokers (in terms of pack-years) had the highest percentage of abnormal overall ECG results (35.0%), followed by participants who never smoked (28.3%), then moderate lifetime cigarette smokers (27.6%). Participants who currently smoked up to 20 cigarettes per day had the highest percentage of abnormal overall ECG results (35.0%), followed by former smokers (32.8%), participants who never smoked (28.3%), and participants who smoked more than 20 cigarettes per day (23.5%). Participants with type B personalities had a higher percentage of abnormal overall ECG results (33.2%) than did participants with type A personalities (27.8%). Participants with a family history of heart disease had a higher prevalence of abnormal overall ECG results than did participants with no family history of heart disease (35.3% versus 24.6%). Diabetic participants had the highest percentage of abnormal overall ECG results (46.7%), followed by participants in the impaired diabetic class (37.0%), then participants classified as normal (26.4%).

The prevalence of right bundle branch block increased significantly with age (p=0.001). Also significantly associated with right bundle branch block were occupation (p=0.040), lifetime cigarette smoking history (p=0.048), and diabetic class (p=0.001). Enlisted flyers had the highest prevalence of right bundle branch block (4.5%), followed by officers (2.6%), then enlisted groundcrew (1.9%). Heavy lifetime cigarette smokers had the highest prevalence of right bundle branch block (3.5%), followed by nonsmokers (2.2%), then moderate lifetime smokers (1.5%). Diabetic participants had the highest percentage of right bundle branch block (5.4%), followed by participants in the impaired diabetic class (2.6%), then participants classified as normal (1.9%).

The percentage of non-specific ST- and T-wave changes increased with age (p=0.001) and body fat (p=0.001), while decreasing with lifetime alcohol use (p=0.024). Family history of heart disease (p=0.001) and diabetic class (p=0.001) also were significant. Participants with a family history of heart disease had a higher percentage of non-specific ST- and T-wave changes than did participants with no history (21.1% versus 14.0%). Diabetic participants had the highest prevalence of non-specific ST- and T-wave changes (29.3%), followed by participants in the impaired diabetic class (24.5%), then participants classified as normal (14.6%).

The prevalence of bradycardia increased significantly with HDL cholesterol levels (p=0.043), while decreasing with the cholesterol-HDL ratio (p=0.005) and body fat (p=0.001). Occupation and diabetic

class also were significantly related to bradycardia (p=0.001 each). Officers had the highest prevalence of bradycardia (5.6%), followed by enlisted flyers (3.0%), then enlisted groundcrew (1.8%). Participants in the normal diabetic class had the highest prevalence of bradycardia (4.5%), followed by diabetic participants (1.7%), then participants in the impaired diabetic class (0.4%).

Tachycardia was significantly associated with lifetime alcohol history (p=0.029) and diabetic class (p=0.008). Non-drinkers had the highest prevalence of tachycardia (1.7%), followed by heavy drinkers (0.8%), then moderate lifetime alcohol drinkers (0.2%). Diabetic participants had the highest prevalence of tachycardia (1.4%), followed by participants in the impaired diabetic class (0.4%), then participants classified as normal (0.2%).

The percentage of participants with arrhythmia increased with age (p=0.001).

Evidence of prior myocardial infarction from the ECG increased with age (p=0.001) and decreased with cholesterol levels (p=0.007). Lifetime cigarette smoking history (p=0.003) and diabetic class (p=0.001) also were significantly associated with prior myocardial infarction. Heavy lifetime cigarette smokers had the highest prevalence of a prior myocardial infarction (5.8%), followed by nonsmokers (2.9%), then moderate lifetime cigarette smokers (2.7%). Diabetic participants had the highest percentage of participants with evidence of a prior myocardial infarction (9.4%), followed by participants in the impaired diabetic class (5.1%), then participants classified as normal (2.8%).

The prevalence of abnormal funduscopic examination results increased with age (p=0.001), lifetime cigarette smoking history (p=0.001), and body fat (p=0.004). Occupation (p=0.001), current cigarette smoking (p=0.019), personality type (p=0.001), and diabetic class (p=0.001) were also significantly associated with an abnormal funduscopic examination. Enlisted flyers had the highest percentage of abnormal funduscopic examination results (18.6%), followed by enlisted groundcrew (11.5%), then officers (11.1%). Participants who never smoked had the lowest percentage of abnormal funduscopic exam results (8.9%), followed by participants who currently smoked up to 20 cigarettes per day (13.5%), former smokers (14.0%), and participants who currently smoked more than 20 cigarettes per day (14.1%). Abnormal funduscopic examinations were more prevalent for participants with personality type B than those with personality type A (14.4% versus 9.2%). Diabetic participants had the highest percentage of abnormal funduscopic exam results (20.0%), followed by participants in the impaired diabetic class (14.3%), then participants classified as normal (10.3%).

The percentage of participants with carotid bruits present increased with age (p=0.001) and lifetime cigarette smoking history (p=0.003). Current cigarette smoking and diabetic class also were significantly associated with carotid bruits (p=0.023 and p=0.007, respectively). Participants who currently smoked up to 20 cigarettes per day had the highest percentage of carotid bruits present (4.1%), followed by participants who currently smoked more than 20 cigarettes per day (3.7%), former smokers (3.1%), and participants who never smoked (1.0%). Diabetic participants had the highest prevalence of carotid bruits (5.1%), followed by participants in the impaired diabetic class (2.9%), then participants classified as normal (2.1%).

Tests of covariate association showed race (p=0.018), lifetime alcohol history (p=0.006), current alcohol use (p=0.005), and current cigarette smoking (p=0.010) to be significantly associated with abnormal radial pulses. The prevalence of abnormal results increased with lifetime alcohol use. Black participants had a higher percentage of abnormal radial pulses than non-Blacks (2.4% versus 0.4%). Participants who currently were moderate drinkers (in terms of drinks per day) had the highest percentage of abnormal radial pulses (1.6%), followed by light drinkers (0.3%), then participants who were the heaviest drinkers (0.0%). Participants who currently smoked up to 20 cigarettes per day had the highest percentage of

abnormal radial pulses (1.9%), followed by participants who currently smoked more than 20 cigarettes per day (0.7%), former smokers (0.4%), and participants who never smoked (0.2%).

The prevalence of abnormal femoral pulses increased with age (p=0.009), lifetime alcohol history (p=0.002), and lifetime cigarette smoking history (p=0.002). Also significant were current alcohol use (p=0.001), current cigarette smoking (p=0.001), and diabetic class (p=0.003). Participants who were currently moderate drinkers had the highest percentage of abnormal femoral pulses (4.4%), followed by the heaviest drinkers (4.0%), then the light drinkers (1.0%). Participants who currently smoked up to 20 cigarettes per day had the highest percentage of abnormal femoral pulses (4.9%), followed by participants who currently smoked more than 20 cigarettes per day (4.4%), former smokers (1.2%), and participants who never smoked (0.3%). Diabetic participants had the highest percentage of abnormal femoral pulses (3.7%), followed by participants classified as normal (1.2%), then participants in the impaired diabetic class (1.1%).

The percentage of participants with abnormal popliteal pulses increased with age (p=0.001), lifetime alcohol history (p=0.013), current alcohol use (p=0.002), lifetime cigarette smoking history (p=0.001), and current cigarette smoking (p=0.001). The association with diabetic class also was significant (p=0.001). Participants who were currently moderate drinkers had the highest percentage of abnormal popliteal pulses (4.9%), followed by the heaviest drinkers (4.0%), then participants who were the lightest drinkers (1.9%). Participants who currently smoked up to 20 cigarettes per day had the highest percentage of abnormal popliteal pulses (7.1%), followed by participants who currently smoked more than 20 cigarettes per day (5.1%), former smokers (2.0%), and participants who never smoked (0.5%). Diabetic participants had the highest percentage of abnormal popliteal pulses (6.0%), followed by participants in the impaired diabetic class (1.8%), then participants classified as normal (1.7%).

The prevalence of abnormal dorsalis pedis pulses increased with age (p=0.001), lifetime cigarette smoking history (p=0.001), and current cigarette smoking (p=0.001). Lifetime alcohol history and diabetic class also were significant (p=0.009 and p=0.001, respectively). Heavy lifetime alcohol drinkers had the highest percentage of abnormal dorsalis pedis pulses (10.6%), followed by non-drinkers (8.5%), then moderate lifetime alcohol drinkers (6.6%). Diabetic participants had the highest prevalence of abnormal dorsalis pedis pulses (14.0%), followed by participants classified as normal (6.7%), then participants in the impaired diabetic class (5.5%).

The percentage of abnormal posterior tibial pulses increased with age (p=0.001), lifetime alcohol history (p=0.027), current alcohol use (p=0.003), lifetime cigarette smoking history (p=0.001), and current cigarette smoking (p=0.001). Personality type and diabetic class also were significantly associated with posterior tibial pulses (p=0.020 and p=0.001, respectively). Participants with type B personalities had more abnormal posterior tibial pulses than participants with type A personalities (6.7% versus 4.2%). Diabetic participants had the highest prevalence of abnormal posterior tibial pulses (13.4%), followed by participants in the impaired diabetic class (5.5%), then participants classified as normal (4.1%).

Abnormal leg pulses increased with age (p=0.001), lifetime cigarette smoking history (p=0.001), and current cigarette smoking (p=0.001). Occupation (p=0.044), lifetime alcohol history (p=0.013), and personality type (p=0.012) also were associated significantly with leg pulses. Enlisted flyers had the highest percentage of abnormal leg pulses (14.2%), followed by enlisted groundcrew (10.0%), then officers (9.3%). Heavy lifetime alcohol drinkers had the highest percentage of abnormal leg pulses (13.4%), followed by non-drinkers (11.0%), then moderate lifetime alcohol drinkers (9.0%). Participants with type B personalities had more abnormal leg pulses than participants with type A personalities (11.7% versus 8.2%). Diabetic participants had the highest prevalence of abnormal leg pulses (18.8%),

followed by participants classified as normal (8.7%), then participants in the impaired diabetic class (8.4%).

The prevalence of abnormal peripheral pulses increased with age (p=0.001), lifetime cigarette smoking history (p=0.001), and current cigarette smoking (p=0.001), while decreasing with body fat (p=0.034). Lifetime alcohol history (p=0.005), current alcohol use (p=0.036), personality type (p=0.026), and diabetic class (p=0.001) also were associated significantly with abnormal peripheral pulses. Heavy lifetime alcohol drinkers had the highest percentage of abnormal peripheral pulses (14.0%), followed by non-drinkers (11.0%) and moderate lifetime alcohol drinkers (9.1%). Participants who were currently moderate drinkers had the highest percentage of abnormal peripheral pulses (14.2%), followed by the heaviest drinkers (14.0%), then participants who were the lightest drinkers (9.8%). Participants with type B personalities had a higher percentage of abnormal peripheral pulses than did participants with type A personalities (11.8% versus 8.7%). Diabetic participants had the highest prevalence of abnormal peripheral pulses (19.4%), followed by participants classified as normal (8.9%), then participants in the impaired diabetic class (8.4%).

The percentage of abnormal intermittent claudication and vascular insufficiency index (ICVI) results increased with lifetime cigarette smoking (p=0.001) and current cigarette smoking (p=0.001). Diabetic class was also significant (p=0.001). Diabetic participants had the highest percentage of abnormal ICVI results (9.1%), followed by participants in the impaired diabetic class (2.9%), then participants classified as normal (2.6%).

14.2.2 Exposure Analysis

The following section presents results of the statistical analysis of the dependent variables shown in Table 14-1. Dependent variables were derived from a medical records review and verification, physical examination and ECG determinations, and an ICVI index based on participant responses to three questions regarding leg pain.

Four models were examined for each dependent variable given in Table 14-1. The analyses of these models are presented below. Further details on dioxin and the modeling strategy are found in Chapters 2 and 7, respectively. These analyses were performed both unadjusted and adjusted for relevant covariates. Model 1 examined the relation between the dependent variable and group (i.e., Ranch Hand or Comparison). In this model, exposure was defined as "yes" for Ranch Hands and "no" for Comparisons without regard to the magnitude of the exposure. As an attempt to quantify exposure, three contrasts of Ranch Hands and Comparisons were performed along with the overall Ranch Hand versus Comparison contrast. These three contrasts compared Ranch Hands and Comparisons within each occupational category (i.e., officers, enlisted flyers, and enlisted groundcrew). As described in previous reports and Table 2-8, the average levels of exposure to dioxin were highest for enlisted groundcrew, followed by enlisted flyers, then officers.

Model 2 explored the relation between the dependent variable and an extrapolated initial dioxin measure for Ranch Hands who had a 1987 dioxin measurement greater than 10 ppt. If a participant did not have a 1987 dioxin level, the 1992 level was used to estimate the initial dioxin level. If a participant did not have a 1987 or a 1992 dioxin level, the 1997 level was used to estimate the initial dioxin level. A statistical adjustment for the percentage of body fat at the time of the participant's blood measurement of dioxin was included in this model to account for body-fat-related differences in elimination rate (52).

Model 3 divided the Ranch Hands examined in Model 2 into two categories based on their initial dioxin measures. These two categories are referred to as "low Ranch Hand" and "high Ranch Hand." Two

additional categories, Ranch Hands with 1987 serum dioxin levels at or below 10 ppt and Comparisons with 1987 serum dioxin levels at or below 10 ppt, were formed and included in the model. Ranch Hands with 1987 serum dioxin levels at or below 10 ppt are referred to as the "background Ranch Hand" category. Dioxin levels in 1992 were used if the 1987 level was not available, and dioxin levels in 1997 were used if the 1987 and 1992 levels were not available. These four categories—Comparisons, background Ranch Hands, low Ranch Hands, and high Ranch Hands—were used in Model 3 analyses. The relation between the dependent variable in each of the three Ranch Hand categories and the dependent variable in the Comparison category was examined. A fourth contrast, exploring the relation of the dependent variable in the combined low and high Ranch Hand categories relative to Comparisons, also was conducted. This combination is referred to in the tables as the "low plus high Ranch Hand" category. As in Model 2, a statistical adjustment for the percentage of body fat at the time of the participant's blood measurement of dioxin was included in this model.

Model 4 examined the relation between the dependent variable and 1987 lipid-adjusted dioxin levels in all Ranch Hands with a dioxin measurement. If a participant did not have a 1987 dioxin measurement, the 1992 measurement was used to determine the dioxin level. If a participant did not have a 1987 or a 1992 dioxin measurement, the 1997 measurement was used to determine the dioxin level.

14.2.2.1 Medical Records Variables

14.2.2.1.1 Essential Hypertension

All Model 1, 2, and 3 analyses of essential hypertension revealed no significant results (Table 14-3(a–f): p>0.13 for each analysis).

The unadjusted and adjusted Model 4 analyses each showed significant positive associations between essential hypertension and 1987 dioxin (Table 14-3(g,h): Est. RR=1.22, p<0.001; Adj. RR=1.18, p=0.011). The percentages of participants with essential hypertension in the low, medium, and high 1987 dioxin categories were 34.0, 38.0, and 49.1, respectively.

Table 14-3. Analysis of Essential Hypertension

Occupational Category	Group	n	Number (%) Yes	Est. Relative Risk (95% C.I.)	p-Value
All	Ranch Hand Comparison	850 1,220	345 (40.6) 509 (41.7)	0.95 (0.80,1.14)	0.606
Officer	Ranch Hand Comparison	329 480	128 (38.9) 199 (41.5)	0.90 (0.68,1.20)	0.467
Enlisted Flyer	Ranch Hand Comparison	149 184	71 (47.7) 80 (43.5)	1.18 (0.77,1.83)	0.447
Enlisted Groundcrew	Ranch Hand Comparison	372 556	146 (39.2) 230 (41.4)	0.92 (0.70,1.20)	0.519

Table 14-3. Analysis of Essential Hypertension (Continued)

(b) MODEL 1: RANCH HANDS VS. COMPARISONS – ADJUSTED

(b) NODEL 1. REPORTED VO. COMPTRICONO PROCESTED							
Adjusted Relative Risk Occupational Category (95% C.I.) p-Value							
All	0.96 (0.79,1.17)	0.708					
Officer	0.85 (0.63,1.16)	0.317					
Enlisted Flyer	1.27 (0.79,2.04)	0.316					
Enlisted Groundcrew	0.96 (0.72,1.29)	0.811					

(c) MODEL 2: RANCH HANDS – INITIAL DIOXIN – UNADJUSTED

Initial 1	Dioxin Category S	Summary Statistics	Analysis Results for Log ₂ (Initial Dioxin) ^a		
Initial Dioxin n		Number (%) Yes	Estimated Relative Risk (95% C.I.) ^b	p-Value		
Low	152	65 (42.8)	1.06 (0.91,1.23)	0.441		
Medium	160	72 (45.0)				
High	159	77 (48.4)				

^a Adjusted for percent body fat at the time of the blood measurement of dioxin. ^b Relative risk for a twofold increase in initial dioxin.

Note: Low = 27-63 ppt; Medium = >63-152 ppt; High = >152 ppt.

(d) MODEL 2: RANCH HANDS - INITIAL DIOXIN - ADJUSTED

Analysis Results for Log ₂ (Initial Dioxin)				
Adjusted Relative Risk				
(95% C.I.) ^a	p-Value			
1.10 (0.91,1.32)	0.314			
	Adjusted Relative Risk (95% C.I.) ^a			

^a Relative risk for a twofold increase in initial dioxin.

(e) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY – UNADJUSTED

		Number (%)	Est. Relative Risk	
Dioxin Category	n	Yes	(95% C.I.) ^{ab}	p-Value
Comparison	1,183	490 (41.4)		
Background RH	372	127 (34.1)	0.86 (0.67,1.11)	0.246
Low RH	229	94 (41.0)	0.95 (0.71,1.29)	0.758
High RH	242	120 (49.6)	1.22 (0.91,1.63)	0.177
Low plus High RH	471	214 (45.4)	1.08 (0.87,1.35)	0.488

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: $1987 \text{ Dioxin} \le 10 \text{ ppt.}$

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, $10 \text{ ppt} < \text{Initial Dioxin} \le 94 \text{ ppt}$.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

^b Adjusted for percent body fat at the time of the blood measurement of dioxin.

Table 14-3. Analysis of Essential Hypertension (Continued)

(f) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY - ADJUSTED

	Adjusted Relative Risk		
Dioxin Category	n	(95% C.I.) ^a	p-Value
Comparison	1,145		
Background RH	356	0.87 (0.66,1.14)	0.320
Low RH	217	0.87 (0.63,1.20)	0.395
High RH	235	1.27 (0.93,1.74)	0.131
Low plus High RH	452	1.06 (0.84,1.35)	0.624

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: $1987 \text{ Dioxin} \le 10 \text{ ppt.}$

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, $10 \text{ ppt} < \text{Initial Dioxin} \le 94 \text{ ppt}$. High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

(g) MODEL 4.	RANCH HANDS -	- 1987 DIOXIN -	- UNADJUSTED
(2) MIODED T.	MAINCH HANDS	1707 DIOMIN	UNADJUBILD

1987 Dioxin Category Summary Statistics		Analysis Results for Log ₂	(1987 Dioxin + 1)	
1987 Dioxin	n	Number (%) Yes	Estimated Relative Risk (95% C.I.) ^a	p-Value
Low	282	96 (34.0)	1.22 (1.11,1.34)	< 0.001
Medium	276	105 (38.0)		
High	285	140 (49.1)		

^a Relative risk for a twofold increase in 1987 dioxin.

Note: Low = ≤ 7.9 ppt; Medium = > 7.9 - 19.6 ppt; High = > 19.6 ppt.

(h) MODEL 4: RANCH HANDS	(h) MODEL 4: RANCH HANDS – 1987 DIOXIN – ADJUSTED				
	Analysis Results for Log ₂ (1987 Dioxin + 1)				
	Adjusted Relative Risk				
n	(95% C.I.) ^a	p-Value			
808	1.18 (1.04,1.34)	0.011			

^a Relative risk for a twofold increase in 1987 dioxin.

14.2.2.1.2 Heart Disease (Excluding Essential Hypertension)

The unadjusted and adjusted Model 1 analyses of a history of heart disease each showed significant group differences when combining all occupations (Table 14-4(a,b): Est. RR=1.26, p=0.013; Adj. RR=1.26, p=0.018, respectively). The percentage of Ranch Hands with heart disease was 66.1 versus 60.8 percent for Comparisons. Stratifying by occupation, unadjusted and adjusted analyses revealed group differences within the enlisted flyer stratum (Table 14-4(a,b): Est. RR=2.10, p=0.003; Adj. RR=2.05; p=0.004, respectively). The percentage of Ranch Hand enlisted flyers with heart disease was 75.2 versus 59.7 percent for the Comparison enlisted flyers.

Table 14-4. Analysis of Heart Disease (Excluding Essential Hypertension)

(a) MODEL 1:	RANCH HANDS VS.	COMPARISONS -	UNADJUSTED

Occupational Category	Group	n	Number (%) Yes	Est. Relative Risk (95% C.I.)	p-Value
All	Ranch Hand Comparison	859 1,232	568 (66.1) 749 (60.8)	1.26 (1.05,1.51)	0.013
Officer	Ranch Hand Comparison	334 484	238 (71.3) 324 (66.9)	1.22 (0.90,1.66)	0.191
Enlisted Flyer	Ranch Hand Comparison	149 186	112 (75.2) 111 (59.7)	2.10 (1.27,3.28)	0.003
Enlisted Groundcrew	Ranch Hand Comparison	376 562	218 (58.0) 314 (55.9)	1.10 (0.84,1.42)	0.523

(b) MODEL 1: RANCH HANDS VS. COMPARISONS – ADJUSTED

	Adjusted Relative Risk	
Occupational Category	(95% C.I.)	p-Value
All	1.26 (1.04,1.53)	0.018
Officer	1.21 (0.88,1.66)	0.238
Enlisted Flyer	2.10 (1.28,3.45)	0.004
Enlisted Groundcrew	1.10 (0.83,1.46)	0.496

(c) MODEL 2: RANCH HANDS – INITIAL DIOXIN – UNADJUSTED

Initial Dioxin Category Summary Statistics		Analysis Results for Log ₂ (Initial Dioxin) ^a	
Initial Dioxin	n	Number (%) Yes	Estimated Relative Risk (95% C.I.) ^b	p-Value
Low	155	115 (74.2)	0.79 (0.68,0.91)	0.001
Medium	161	99 (61.5)		
High	160	88 (55.0)		

^a Adjusted for percent body fat at the time of the blood measurement of dioxin. ^b Relative risk for a twofold increase in initial dioxin.

Note: Low = 27-63 ppt; Medium = >63-152 ppt; High = >152 ppt.

(d) MODEL 2: RANCH I	HANDS – INITIAL DIOXIN – ADJUSTED	
	Analysis Results for Log ₂ (Initial Dioxi	n)
	Adjusted Relative Risk	
n	(95% C.I.) ^a	p-Value
457	0.90 (0.75,1.08)	0.249

^a Relative risk for a twofold increase in initial dioxin.

Table 14-4. Analysis of Heart Disease (Excluding Essential Hypertension) (Continued)

(e) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY – UNADJUSTED

Di i G		Number (%)	Est. Relative Risk	*** 1
Dioxin Category	n	Yes	(95% C.I.) ^{ab}	p-Value
Comparison	1,195	730 (61.1)		
Background RH	376	259 (68.9)	1.43 (1.11,1.83)	0.005
Low RH	233	163 (70.0)	1.48 (1.09,2.00)	0.011
High RH	243	139 (57.2)	0.84 (0.64,1.11)	0.228
Low plus High RH	476	302 (63.4)	1.11 (0.89,1.39)	0.359

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: 1987 Dioxin ≤ 10 ppt.

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, $10 \text{ ppt} < \text{Initial Dioxin} \le 94 \text{ ppt}$.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

(f) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY - ADJUSTED

		Adjusted Relative Risk	
Dioxin Category	n	(95% C.I.) ^a	p-Value
Comparison	1,155		
Background RH	360	1.34 (1.03,1.75)	0.032
Low RH	221	1.33 (0.96,1.84)	0.081
High RH	236	1.03 (0.76,1.40)	0.865
Low plus High RH	457	1.16 (0.92,1.48)	0.209

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: $1987 \text{ Dioxin} \le 10 \text{ ppt.}$

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, $10 \text{ ppt} < \text{Initial Dioxin} \le 94 \text{ ppt}$.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

(g) MODEL 4: RANCH HANDS – 1987 DIOXIN – UNADJUSTED

1987 Dioxin Category Summary Statistics		Analysis Results for Log ₂ (1987 Dioxin + 1)		
1987 Number (%) Dioxin n Yes		Estimated Relative Risk (95% C.I.) ^a	p-Value	
Low	284	192 (67.6)	0.87 (0.79,0.96)	0.004
Medium	281	199 (70.8)		
High	287	170 (59.2)		

^a Relative risk for a twofold increase in 1987 dioxin.

Note: Low = \le 7.9 ppt; Medium = >7.9–19.6 ppt; High = >19.6 ppt.

^b Adjusted for percent body fat at the time of the blood measurement of dioxin.

Table 14-4. Analysis of Heart Disease (Excluding Essential Hypertension) (Continued)

(h) MODEL 4: RANCH HANDS – 1987 DIOXIN – ADJUSTED							
	Analysis Results for Log_2 (1987 Dioxin + 1)						
	Adjusted Relative Risk						
n	(95% C.I.) ^a	p-Value					
817	0.92 (0.81,1.04)	0.159					

^a Relative risk for a twofold increase in 1987 dioxin.

The unadjusted Model 2 analysis revealed a significant inverse association between heart disease and initial dioxin (Table 14-4(c): Est. RR=0.79, p=0.001). The percentages of participants with heart disease in the low, medium, and high initial dioxin categories were 74.2, 61.5, and 55.0, respectively. After covariate adjustment, the results became nonsignificant (Table 14-4(d): p=0.249).

The Model 3 unadjusted analysis of heart disease revealed two significant contrasts: Ranch Hands in the background dioxin category versus Comparisons and Ranch Hands in the low dioxin category versus Comparisons (Table 14-4(e): Est. RR=1.43, p=0.005; Est. RR=1.48, p=0.011, respectively). The adjusted analysis showed a significant difference between Ranch Hands in the background dioxin category and Comparisons (Table 14-4(f): Adj. RR=1.34, p=0.032) and a marginally significant difference between Ranch Hands in the low dioxin category and Comparisons (Table 14-4(f): Adj. RR=1.33, p=0.081). The percentages of participants with heart disease for Ranch Hands in the background dioxin category, Ranch Hands in the low dioxin category, and Comparisons were 68.9, 70.0, and 61.1, respectively.

The Model 4 unadjusted analysis showed a significant inverse association between heart disease and 1987 dioxin (Table 14-4(g): Est. RR=0.87, p=0.004). The percentages of participants with heart disease in the low, medium, and high 1987 dioxin categories were 67.6, 70.8, and 59.2, respectively. The results became nonsignificant after adjusting for covariates (Table 14-4(h): p=0.159).

14.2.2.1.3 Myocardial Infarction

All unadjusted and adjusted Model 1 through Model 4 analyses of myocardial infarction were nonsignificant (Table 14-5(a–h): p>0.10 for each analysis).

Table 14-5. Analysis of Myocardial Infarction

(a) MODEL 1: RANCH HANDS VS. COMPARISONS – UNADJUSTED						
Occupational Category	Group	n	Number (%) Yes	Est. Relative Risk (95% C.I.)	p-Value	
All	Ranch Hand Comparison	859 1,232	74 (8.6) 102 (8.3)	1.04 (0.76,1.43)	0.786	
Officer	Ranch Hand Comparison	334 484	28 (8.4) 42 (8.7)	0.96 (0.58,1.59)	0.882	
Enlisted Flyer	Ranch Hand Comparison	149 186	16 (10.7) 15 (8.1)	1.37 (0.65,2.87)	0.403	
Enlisted Groundcrew	Ranch Hand Comparison	376 562	30 (8.0) 45 (8.0)	1.00 (0.62,1.61)	0.987	

Table 14-5. Analysis of Myocardial Infarction (Continued)

(b) MODEL 1: RANCH HANDS VS. COMPARISONS – ADJUSTED

(b) MODEL 1: RANCH HANDS VS	(D) MODEL 1: RANCH HANDS VS. COMPARISONS - ADJUSTED					
Occupational Category	Adjusted Relative Risk (95% C.I.)	p-Value				
All	1.02 (0.73,1.42)	0.915				
Officer	0.86 (0.50,1.46)	0.567				
Enlisted Flyer	1.57 (0.72,3.43)	0.255				
Enlisted Groundcrew	0.99 (0.59,1.67)	0.975				

(c) MODEL 2: RANCH HANDS – INITIAL DIOXIN – UNADJUSTED

Initial Dioxin Category Summary Statistics		Analysis Results for Log ₂ (Initial Dioxin) ^a	
Initial Dioxin	n	Number (%) Yes	Estimated Relative Risk (95% C.I.) ^b	p-Value
Low	155	12 (7.7)	1.01 (0.79,1.28)	0.945
Medium	161	18 (11.2)		
High	160	13 (8.1)		

^a Adjusted for percent body fat at the time of the blood measurement of dioxin.

Note: Low = 27-63 ppt; Medium = >63-152 ppt; High = >152 ppt.

(d) MODEL 2: RANCH HANDS – INITIAL DIOXIN – ADJUSTED

Analysis Results for Log ₂ (Initial Dioxin)					
Adjusted Rel					
n (95% C	(A.I.) ^a p-Value				
457 1.30 (0.95	5,1.77) 0.106				

^a Relative risk for a twofold increase in initial dioxin.

Note: Results are not adjusted for race because of the sparse number of participants with a myocardial infarction.

(e) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY – UNADJUSTED

		Number (%)	Est. Relative Risk	
Dioxin Category	n	Yes	(95% C.I.) ^{ab}	p-Value
Comparison	1,195	98 (8.2)		
Background RH	376	29 (7.7)	0.98 (0.63,1.51)	0.919
Low RH	233	19 (8.2)	0.99 (0.59,1.65)	0.958
High RH	243	24 (9.9)	1.18 (0.73,1.89)	0.496
Low plus High RH	476	43 (9.0)	1.08 (0.74,1.58)	0.689

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: $1987 \text{ Dioxin} \le 10 \text{ ppt.}$

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, 10 ppt < Initial Dioxin ≤ 94 ppt. High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

^b Relative risk for a twofold increase in initial dioxin.

^b Adjusted for percent body fat at the time of the blood measurement of dioxin.

Table 14-5. Analysis of Myocardial Infarction (Continued)

(f) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY - ADJUSTED

		Adjusted Relative Risk	
Dioxin Category	n	(95% C.I.) ^a	p-Value
Comparison	1,155		
Background RH	360	0.89 (0.55,1.43)	0.625
Low RH	221	0.84 (0.49,1.46)	0.544
High RH	236	1.39 (0.83,2.32)	0.215
Low plus High RH	457	1.09 (0.73,1.63)	0.673

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: 1987 Dioxin ≤ 10 ppt.

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, 10 ppt < Initial Dioxin ≤ 94 ppt. High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

(g) MODEL 4: RANCH HANDS – 1987 DIOXIN – UNADJUSTED

(g) MODEL II		150. 210111	CIMIDGESTED	
1987 Dioxin Category Summary Statistics		Analysis Results for Log ₂	(1987 Dioxin + 1)	
1987 Number (%) Dioxin n Yes		Estimated Relative Risk (95% C.I.) ^a	p-Value	
Low	284	21 (7.4)	1.03 (0.87,1.21)	0.740
Medium	281	23 (8.2)		
High	287	28 (9.8)		

^a Relative risk for a twofold increase in 1987 dioxin.

Note: Low = \le 7.9 ppt; Medium = >7.9–19.6 ppt; High = >19.6 ppt.

(h) MODEL 4: RANCH HANDS – 1987 DIOXIN – ADJUSTED					
	Analysis Results for Log ₂ (1987 Dioxin + 1)				
	Adjusted Relative Risk				
n	(95% C.I.) ^a	p-Value			
817	1.16 (0.94,1.44)	0.170			

^a Relative risk for a twofold increase in 1987 dioxin.

14.2.2.1.4 Stroke or Transient Ischemic Attack

All analysis results of stroke or transient ischemic attack were nonsignificant (Table 14-6(a-h): $p \ge 0.10$ for each analysis).

Table 14-6. Analysis of Stroke or Transient Ischemic Attack

(a) MODEL 1: RANCH HANDS VS. COMPARISONS – UNADJUSTED						
Occupational Category	Group	n	Number (%) Yes	Est. Relative Risk (95% C.I.)	p-Value	
All	Ranch Hand Comparison	859 1,232	11 (1.3) 14 (1.1)	1.13 (0.51,2.50)	0.766	
Officer	Ranch Hand	334	5 (1.5)	1.46 (0.42,5.07)	0.555	

Officer	Ranch Hand	334	5 (1.5)	1.46 (0.42,5.07)	0.555
	Comparison	484	5 (1.0)		
Enlisted Flyer	Ranch Hand	149	0 (0.0)		0.330^{a}
	Comparison	186	3 (1.6)		
Enlisted	Ranch Hand	376	6 (1.6)	1.50 (0.48,4.69)	0.483
Groundcrew	Comparison	562	6 (1.1)		

^a P-value determined using a chi-square test with continuity correction because of the sparse number of participants with a stroke or transient ischemic attack.

^{--:} Results not presented because of the sparse number of participants with a stroke or transient ischemic attack.

(b) MODEL 1: RANCH HANDS VS. COMPARISONS – ADJUSTED							
Adjusted Relative Risk Occupational Category (95% C.I.) p-Value							
All	1.21 (0.51,2.85)	0.666					
Officer Enlisted Flyer Enlisted Groundcrew	1.18 (0.31,4.51) 1.80 (0.53,6.06)	0.806 0.345					

^{--:} Results not presented because of the sparse number of participants with a stroke or transient ischemic attack.

(c) MODEL 2: RANCH HANDS – INITIAL DIOXIN – UNADJUSTED							
Initial 1	Initial Dioxin Category Summary Statistics Analysis Results for Log ₂ (Initial Dioxin) ^a						
Initial Dioxin	n	Number (%) Yes	Estimated Relative Risk (95% C.I.) ^b	p-Value			
Low	155	1 (0.6)	1.22 (0.68,2.16)	0.513			
Medium	161	2 (1.2)					
High	160	3 (1.9)					

^a Adjusted for percent body fat at the time of the blood measurement of dioxin.

Note: Low = 27-63 ppt; Medium = >63-152 ppt; High = >152 ppt.

^b Relative risk for a twofold increase in initial dioxin.

Table 14-6. Analysis of Stroke or Transient Ischemic Attack (Continued)

(d) MODEL 2: RANCH HANDS – INITIAL DIOXIN – ADJUSTED

(d) MODEL 2: RANCH H	IANDS – INITIAL DIOXIN – ADJUSTED	
	Analysis Results for Log ₂ (Initial Diox	in)
	Adjusted Relative Risk	
n	(95% C.I.) ^a	p-Value
457	1.33 (0.72,2.47)	0.379

^a Relative risk for a twofold increase in initial dioxin.

Note: Results are not adjusted for race and occupation because of the sparse number of Ranch Hands with a stroke or transient ischemic attack.

(e) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY - UNADJUSTED

		Number (%)	Est. Relative Risk	
Dioxin Category	n	Yes	(95% C.I.) ^{ab}	p-Value
Comparison	1,195	14 (1.2)		
Background RH	376	5 (1.3)	1.13 (0.40,3.18)	0.816
Low RH	233	1 (0.4)	0.36 (0.05,2.78)	0.330
High RH	243	5 (2.1)	1.78 (0.63,5.02)	0.275
Low plus High RH	476	6 (1.3)	0.82 (0.25,2.68)	0.741

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: 1987 Dioxin ≤ 10 ppt.

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, 10 ppt < Initial Dioxin ≤ 94 ppt.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

(f) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY - ADJUSTED

		Adjusted Relative Risk	
Dioxin Category	n	(95% C.I.) ^a	p-Value
Comparison	1,155		
Background RH	360	0.97 (0.30,3.16)	0.956
Low RH	221	0.42 (0.05,3.26)	0.404
High RH	236	2.65 (0.83,8.46)	0.100
Low plus High RH	457	1.08 (0.32,3.71)	0.900

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: $1987 \text{ Dioxin} \le 10 \text{ ppt.}$

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, 10 ppt < Initial Dioxin ≤ 94 ppt.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

^b Adjusted for percent body fat at the time of the blood measurement of dioxin.

Table 14-6. Analysis of Stroke or Transient Ischemic Attack (Continued)

(g) MODEL 4: RANCH HANDS – 1987 DIOXIN – UNADJUSTED						
1987 Dioxin	(1987 Dioxin + 1)					
1987 Dioxin	n	Number (%) Yes	Estimated Relative Risk (95% C.I.) ^a	p-Value		
Low	284	4 (1.4)	0.99 (0.66,1.48)	0.957		
Medium	281	2 (0.7)				
High	287	5 (1.7)				

^a Relative risk for a twofold increase in 1987 dioxin.

Note: Low = ≤ 7.9 ppt; Medium = > 7.9 - 19.6 ppt; High = > 19.6 ppt.

(h) MODEL 4: RANCH HANDS – 1987 DIOXIN – ADJUSTED							
Analysis Results for Log ₂ (1987 Dioxin + 1)							
	Adjusted Relative Risk						
n	(95% C.I.) ^a	p-Value					
817	1.15 (0.71,1.85)	0.578					

^a Relative risk for a twofold increase in 1987 dioxin.

Note: Results are not adjusted for occupation because of the sparse number of Ranch Hands with a stroke or transient ischemic attack.

14.2.2.2 Physical Examination Variables – Central Cardiac Function

14.2.2.2.1 Systolic Blood Pressure (Continuous)

All Model 1 and Model 2 analyses of systolic blood pressure in its continuous form showed no significant results (Table 14-7(a–d): p>0.23 for each analysis).

Table 14-7. Analysis of Systolic Blood Pressure (Continuous)

Occupational Category	Group	n	Mean ^a	Difference of Means (95% C.I.) ^b	p-Value ^c
All	Ranch Hand Comparison	859 1,232	124.9 125.6	-0.7	0.383
Officer	Ranch Hand Comparison	334 484	125.9 126.2	-0.2	0.865
Enlisted Flyer	Ranch Hand Comparison	149 186	127.0 127.3	-0.3	0.875
Enlisted Groundcrew	Ranch Hand Comparison	376 562	123.1 124.5	-1.4	0.241

^a Transformed from natural logarithm scale.

^b Difference of means after transformation to original scale; confidence interval on difference of means not presented because analysis was performed on natural logarithm scale.

^c P-value is based on difference of means on natural logarithm scale.

Table 14-7. Analysis of Systolic Blood Pressure (Continuous) (Continued)

(b) MODEL 1: RANCH HANDS VS. COMPARISONS – ADJUSTED							
Occupational Category	Group	n	Adjusted Mean ^a	Difference of Adj. Means (95% C.I.) ^b	p-Value ^c		
All	Ranch Hand Comparison	822 1,189	127.7 128.4	-0.6	0.415		
Officer	Ranch Hand Comparison	322 472	127.2 128.1	-0.9	0.468		
Enlisted Flyer	Ranch Hand Comparison	140 178	128.7 128.6	0.1	0.967		
Enlisted Groundcrew	Ranch Hand Comparison	360 539	127.5 128.2	-0.7	0.574		

^a Transformed from natural logarithm scale.

(c) MODEL 2: RANCH HANDS – INITIAL DIOXIN – UNADJUSTED						
Initial Di	Initial Dioxin Category Summary Statistics Analysis Results for Log ₂ (Initial Dioxin) ^b					
Initial Dioxin	n	Mean ^a	Adj. Mean ^{ab}	\mathbb{R}^2	Slope (Std. Error) ^c	p-Value
Low	155	125.8	126.4	0.049	-0.006 (0.005)	0.238
Medium	161	125.7	125.8			
High	160	124.2	123.6			

^a Transformed from natural logarithm scale.

Note: Low = 27-63 ppt; Medium = >63-152 ppt; High = >152 ppt.

(d) MODEL 2: RANCH HANDS – INITIAL DIOXIN – ADJUSTED						
Initial Dioxin	Initial Dioxin Category Summary Statistics Analysis Results for Log ₂ (Initial Dioxin)					
Initial Dioxin	n	Adj. Mean ^a	Adj. Slope R ² (Std. Error) ^b p-Valu			
Low	150	129.0	0.135	-0.000 (0.006)	0.983	
Medium	150	130.2				
High	157	128.5				

^a Transformed from natural logarithm scale.

Note: Low = 27-63 ppt; Medium = >63-152 ppt; High = >152 ppt.

^b Difference of means after transformation to original scale; confidence interval on difference of means not presented because analysis was performed on natural logarithm scale. ^c P-value is based on difference of means on natural logarithm scale.

^b Adjusted for percent body fat at the time of the blood measurement of dioxin.

^c Slope and standard error based on natural logarithm of systolic blood pressure versus log₂ (initial dioxin).

^b Slope and standard error based on natural logarithm of systolic blood pressure versus log₂ (initial dioxin).

Table 14-7. Analysis of Systolic Blood Pressure (Continuous) (Continued)

(e) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY – UNADJUSTED

Dioxin Category	n	Mean ^a	Adj. Mean ^{ab}	Difference of Adj. Mean vs. Comparisons (95% C.I.) ^c	p-Value ^d
	11	Mican	Auj. Mean	(33 /0 C.I.)	p- value
Comparison	1,195	125.6	125.5		
Background RH	376	124.4	125.4	-0.1	0.935
Low RH	233	126.2	125.9	0.4	0.730
High RH	243	124.4	123.4	-2.1	0.079
Low plus High RH	476	125.2	124.6	-0.9	0.346

^a Transformed from natural logarithm scale.

Note: RH = Ranch Hand.

Comparison: 1987 Dioxin ≤ 10 ppt.

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, $10 \text{ ppt} < \text{Initial Dioxin} \le 94 \text{ ppt}$.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

(f) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY - ADJUSTED

Dioxin Category	n	Adj. Mean ^a	Difference of Adj. Mean vs. Comparisons (95% C.I.) ^b	p-Value ^c
Comparison	1,155	128.5		
Background RH	360	128.5	0.0	0.990
Low RH	221	127.9	-0.6	0.651
High RH	236	127.0	-1.5	0.222
Low plus High RH	457	127.4	-1.1	0.262

^a Transformed from natural logarithm scale.

Note: RH = Ranch Hand.

Comparison: 1987 Dioxin ≤ 10 ppt.

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, 10 ppt < Initial Dioxin ≤ 94 ppt.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

^b Adjusted for percent body fat at the time of the blood measurement of dioxin.

^c Difference of means after transformation to original scale; confidence interval on difference of means not presented because analysis was performed on natural logarithm scale.

^d P-value is based on difference of means on natural logarithm scale.

^b Difference of means after transformation to original scale; confidence interval on difference of means not presented because analysis was performed on natural logarithm scale.

^c P-value is based on difference of means on natural logarithm scale.

Table 14-7. Analysis of Systolic Blood Pressure (Continuous) (Continued)

(g) MODEL 4: RANCH HANDS – 1987 DIOXIN – UNADJUSTED					
1987 Dioxin Category Summary Statistics			Analysis Results for Log ₂ (1987 Dioxin +1)		
1987 Dioxin	n	Mean ^a	\mathbb{R}^2	Slope (Std. Error) ^b	p-Value
Low	284	124.0	< 0.001	0.001 (0.003)	0.693
Medium	281	125.9			
High	287	124.8			

^a Transformed from natural logarithm scale.

Note: Low = ≤ 7.9 ppt; Medium = > 7.9 - 19.6 ppt; High = > 19.6 ppt.

(h) MODEL 4: RANCH HANDS – 1987 DIOXIN – ADJUSTED					
1987 Dioxin Category Summary Statistics		Analysis Results for Log ₂ (1987 Dioxin + 1)			
1987 Dioxin	n	Adj. Mean ^a	\mathbb{R}^2	Adjusted Slope (Std. Error) ^b	p-Value
Low Medium	271 271	128.3 127.2	0.126	-0.005 (0.004)	0.165
High	275	127.1			

^a Transformed from natural logarithm scale.

Note: Low = \le 7.9 ppt; Medium = >7.9–19.6 ppt; High = >19.6 ppt.

The unadjusted Model 3 analysis showed a marginally significant difference between Ranch Hands in the high dioxin category and Comparisons (Table 14-7(e): difference of means=-2.1 mm Hg, p=0.079). Ranch Hands in the high dioxin category had a lower mean systolic blood pressure (123.4 mm Hg) than the Comparisons (125.5 mm Hg). The adjusted Model 3 analysis revealed no significant contrasts (Table 14-7(f): p>0.22 for each contrast).

Both the unadjusted and adjusted Model 4 analyses revealed no significant associations between 1987 dioxin and systolic blood pressure in its continuous form (Table 14-7(g,h): p>0.16 for each analysis).

14.2.2.2.2 Systolic Blood Pressure (Discrete)

The unadjusted and adjusted Model 1 analyses of systolic blood pressure in its discrete form showed no significant differences between Ranch Hands and Comparisons when examined across all occupations and within each occupation (Table 14-8(a,b): p>0.63 for each contrast).

^b Slope and standard error based on natural logarithm of systolic blood pressure versus log₂ (1987 dioxin + 1).

^b Slope and standard error based on natural logarithm of systolic blood pressure versus log₂ (1987 dioxin + 1).

Table 14-8. Analysis of Systolic Blood Pressure (Discrete)

(a) MODEL 1: RANCH HANDS VS. COMPARISONS – UNADJUSTED

Occupational Category	Group	n	Number (%) High	Est. Relative Risk (95% C.I.)	p-Value
All	Ranch Hand Comparison	859 1,232	181 (21.1) 262 (21.3)	0.99 (0.80,1.22)	0.914
Officer	Ranch Hand Comparison	334 484	78 (23.4) 112 (23.1)	1.01 (0.73,1.41)	0.944
Enlisted Flyer	Ranch Hand Comparison	149 186	36 (24.2) 43 (23.1)	1.06 (0.64,1.76)	0.823
Enlisted Groundcrew	Ranch Hand Comparison	376 562	67 (17.8) 107 (19.0)	0.92 (0.66,1.29)	0.638

(b) MODEL 1: RANCH HANDS VS. COMPARISONS – ADJUSTED

Adjusted Relative Risk					
Occupational Category	(95% C.I.)	p-Value			
All	0.99 (0.79,1.24)	0.899			
Officer	0.95 (0.67,1.35)	0.784			
Enlisted Flyer	1.13 (0.66,1.93)	0.661			
Enlisted Groundcrew	0.96 (0.67,1.38)	0.832			

(c) MODEL 2: RANCH HANDS – INITIAL DIOXIN – UNADJUSTED

Initial Dioxin Category Summary Statistics			Analysis Results for Log ₂ (Initial Dioxin) ^a		
Initial Dioxin	n	Number (%) High	Estimated Relative Risk (95% C.I.) ^b	p-Value	
Low	155	40 (25.8)	0.83 (0.69,0.99)	0.031	
Medium	161	36 (22.4)			
High	160	29 (18.1)			

^a Adjusted for percent body fat at the time of the blood measurement of dioxin. ^b Relative risk for a twofold increase in initial dioxin.

Note: Low = 27-63 ppt; Medium = >63-152 ppt; High = >152 ppt.

(d) MODEL 2: RANCH HANDS – INITIAL DIOXIN – ADJUSTED			
	Analysis Results for Log ₂ (Initial Dioxi	n)	
Adjusted Relative Risk			
n	(95% C.I.) ^a	p-Value	
457	0.89 (0.71,1.11)	0.296	

^a Relative risk for a twofold increase in initial dioxin.

Table 14-8. Analysis of Systolic Blood Pressure (Discrete) (Continued)

(e) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY - UNADJUSTED

D'an's Catalana		Number (%)	Est. Relative Risk	37-1
Dioxin Category	n	High	(95% C.I.) ^{ab}	p-Value
Comparison	1,195	253 (21.2)		
Background RH	376	74 (19.7)	1.00 (0.75,1.34)	0.998
Low RH	233	59 (25.3)	1.25 (0.90,1.73)	0.188
High RH	243	46 (18.9)	0.80 (0.56,1.14)	0.208
Low plus High RH	476	105 (22.1)	0.99 (0.76,1.29)	0.952

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: 1987 Dioxin ≤ 10 ppt.

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, $10 \text{ ppt} < \text{Initial Dioxin} \le 94 \text{ ppt}$.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

(f) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY - ADJUSTED

		Adjusted Relative Risk	
Dioxin Category	n	(95% C.I.) ^a	p-Value
Comparison	1,155		
Background RH	360	1.00 (0.73,1.37)	0.983
Low RH	221	1.12 (0.79,1.59)	0.532
High RH	236	0.84 (0.57,1.23)	0.365
Low plus High RH	457	0.96 (0.73,1.27)	0.791

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: $1987 \text{ Dioxin} \le 10 \text{ ppt.}$

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, $10 \text{ ppt} < \text{Initial Dioxin} \le 94 \text{ ppt}$.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

(g) MODEL 4: RANCH HANDS – 1987 DIOXIN – UNADJUSTED

1987 Dioxin Category Summary Statistics			Analysis Results for Log ₂ (1987 Dioxin + 1)		
1987 Dioxin	n	Number (%) High	Estimated Relative Risk (95% C.I.) ^a	p-Value	
Low	284	54 (19.0)	1.00 (0.89,1.12)	0.956	
Medium	281	66 (23.5)			
High	287	59 (20.6)			

^a Relative risk for a twofold increase in 1987 dioxin.

Note: Low = ≤ 7.9 ppt; Medium = > 7.9 - 19.6 ppt; High = > 19.6 ppt.

^b Adjusted for percent body fat at the time of the blood measurement of dioxin.

Table 14-8. Analysis of Systolic Blood Pressure (Discrete) (Continued)

(h) MODEL 4: RANCH HANDS – 1987 DIOXIN – ADJUSTED						
Analysis Results for Log ₂ (1987 Dioxin + 1)						
	Adjusted Relative Risk					
n	(95% C.I.) ^a	p-Value				
817	0.88 (0.76,1.02)	0.099				

^a Relative risk for a twofold increase in 1987 dioxin.

A significant inverse association between discrete systolic blood pressure and initial dioxin was found in the unadjusted Model 2 analysis (Table 14-8(c): Est. RR=0.83, p=0.031). After adjusting for covariates, the results became nonsignificant (Table 14-8(d): p=0.296).

The unadjusted and adjusted Model 3 analyses of systolic blood pressure showed no significant contrasts between the Ranch Hand dioxin category and Comparisons (Table 14-8(e,f): p>0.18 for each contrast).

The unadjusted Model 4 results were nonsignificant (Table 14-8(g): p=0.956). After adjusting for covariates, the results became marginally significant (Table 14-8(h): Adj. RR=0.88, p=0.099). The percentages of participants with high discrete systolic blood pressures in the low, medium, and high 1987 dioxin categories were 19.0, 23.5, and 20.6, respectively.

14.2.2.2.3 Diastolic Blood Pressure (Continuous)

All Model 1 and Model 2 analyses of diastolic blood pressure in its continuous form showed no significant results (Table 14-9(a−d): p≥0.19 for each analysis).

The unadjusted Model 3 analysis of continuous diastolic blood pressure revealed a marginally significant difference between Ranch Hands in the high dioxin category and Comparisons (Table 14-9(e): difference of means=1.08 mm Hg, p=0.099). The adjusted results were nonsignificant (Table 14-9(f): p>0.13 for each contrast).

A significant positive association between 1987 dioxin and continuous diastolic blood pressure was found in the unadjusted Model 4 analysis (Table 14-9(g): slope=0.031, p=0.014). The mean diastolic blood pressure in the low, medium, and high 1987 dioxin categories was 73.97 mm Hg, 73.76 mm Hg, and 75.94 mm Hg, respectively. After adjusting for covariates, the results became nonsignificant (Table 14-9(h): p=0.315).

Table 14-9. Analysis of Diastolic Blood Pressure (Continuous)

(a) MODEL 1: RANCH HANDS VS. COMPARISONS – UNADJUSTED

Occupational Category	Group	n	Mean ^a	Difference of Means (95% C.I.) ^b	p-Value ^c
All	Ranch Hand Comparison	859 1,232	74.55 74.61	-0.06	0.883
Officer	Ranch Hand Comparison	334 484	74.17 74.21	-0.04	0.952
Enlisted Flyer	Ranch Hand Comparison	149 186	75.22 75.10	0.12	0.905
Enlisted Groundcrew	Ranch Hand Comparison	376 562	74.63 74.80	-0.17	0.780

^a Transformed from square root scale.

^c P-value is based on difference of means on square root scale.

(b) MODEL 1: RANCH HANDS VS. COMPARISONS – ADJUSTED							
Occupational Category	Group	n	Adjusted Mean ^a	Difference of Adj. Means (95% C.I.) ^b	p-Value ^c		
All	Ranch Hand Comparison	822 1,189	75.68 75.62	0.06	0.889		
Officer	Ranch Hand Comparison	322 472	75.29 75.37	-0.08	0.907		
Enlisted Flyer	Ranch Hand Comparison	140 178	76.47 76.13	0.33	0.752		
Enlisted Groundcrew	Ranch Hand Comparison	360 539	75.37 75.29	0.08	0.898		

^a Transformed from square root scale.

^c P-value is based on difference of means on square root scale.

(c) MODEL 2: RANCH HANDS – INITIAL DIOXIN – UNADJUSTED							
Initial Dioxin Category Summary Statistics				Analy	sis Results for Log ₂ (Ini	tial Dioxin) ^b	
Initial Dioxin	n	Mean ^a	Adj. Mean ^{ab}	\mathbb{R}^2	Slope (Std. Error) ^c	p-Value	
Low	155	74.07	74.24	0.023	0.025 (0.019)	0.190	
Medium	161	75.16	75.17				
High	160	76.07	75.89				

^b Difference of means after transformation to original scale; confidence interval on difference of means not presented because analysis was performed on square root scale.

^b Difference of means after transformation to original scale; confidence interval on difference of means not presented because analysis was performed on square root scale.

 ^a Transformed from square root scale.
 ^b Adjusted for percent body fat at the time of the blood measurement of dioxin.

^c Slope and standard error based on square root of diastolic blood pressure versus log₂ (initial dioxin).

Table 14-9. Analysis of Diastolic Blood Pressure (Continuous) (Continued)

(d) MODEL 2: RANCH HANDS - INITIAL DIOXIN - ADJUSTED

` ′					
Initial Dioxin Category Summary Statistics			Analysis Results for Log ₂ (Initial Dioxin)		
Initial Dioxin	n	Adj. Mean ^a	\mathbb{R}^2	Adj. Slope (Std. Error) ^b	p-Value
Low	150	76.09	0.073	0.019 (0.023)	0.425
Medium	150	77.21			
High	157	77.40			

^a Transformed from square root scale.

Note: Low = 27-63 ppt; Medium = >63-152 ppt; High = >152 ppt.

(e) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY – UNADJUSTED

Dioxin Category	n	Mean ^a	Adj. Mean ^{ab}	Difference of Adj. Mean vs. Comparisons (95% C.I.) ^c	p-Value ^d
Comparison	1,195	74.58	74.57		
Background RH	376	73.87	74.14	-0.43	0.432
Low RH	233	74.26	74.19	-0.38	0.569
High RH	243	75.93	75.65	1.08	0.099
Low plus High RH	476	75.11	74.93	0.36	0.468

^a Transformed from square root scale.

Note: RH = Ranch Hand.

Comparison: 1987 Dioxin ≤ 10 ppt.

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, $10 \text{ ppt} < \text{Initial Dioxin} \le 94 \text{ ppt}$.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

^b Slope and standard error based on square root of diastolic blood pressure versus log₂ (initial dioxin).

^b Adjusted for percent body fat at the time of the blood measurement of dioxin.

^c Difference of means after transformation to original scale; confidence interval on difference of means not presented because analysis was performed on square root scale.

^d P-value is based on difference of means on square root scale.

Table 14-9. Analysis of Diastolic Blood Pressure (Continuous) (Continued)

(f) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY - ADJUSTED

Dioxin Category	n	Adj. Mean ^a	Difference of Adj. Mean vs. Comparisons (95% C.I.) ^b	p-Value ^c
Comparison	1,155	75.67		
Background RH	360	75.56	-0.11	0.844
Low RH	221	75.23	-0.44	0.515
High RH	236	76.69	1.02	0.135
Low plus High RH	457	75.98	0.31	0.544

^a Transformed from square root scale.

Note: RH = Ranch Hand.

Comparison: $1987 \text{ Dioxin} \le 10 \text{ ppt.}$

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, $10 \text{ ppt} < \text{Initial Dioxin} \le 94 \text{ ppt}$.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

(g) MODEL 4: RANCH HANDS – 1987 DIOXIN – UNADJUSTED

1987 Dioxin Category Summary Statistics			Analysis l	Results for Log ₂ (1987 D	Pioxin +1)
1987 Dioxin	n	Mean ^a	\mathbb{R}^2	Slope (Std. Error) ^b	p-Value
Low	284	73.97	0.007	0.031 (0.013)	0.014
Medium	281	73.76			
High	287	75.94			

^a Transformed from square root scale.

Note: Low = ≤ 7.9 ppt; Medium = > 7.9 - 19.6 ppt; High = > 19.6 ppt.

(h) MODEL 4: RANCH HANDS – 1987 DIOXIN – ADJUSTED							
1987 Dioxin Category Summary Statistics			Analysis Re	sults for Log ₂ (1987 Dioxin	+ 1)		
1987 Dioxin	n	Adj. Mean ^a	\mathbb{R}^2	Adjusted Slope (Std. Error) ^b	p-Value		
Low	271	75.59	0.061	0.016 (0.016)	0.315		
Medium	271	75.01					
High	275	77.24					

^a Transformed from square root scale.

Note: Low = \le 7.9 ppt; Medium = >7.9–19.6 ppt; High = >19.6 ppt.

^b Difference of means after transformation to original scale; confidence interval on difference of means not presented because analysis was performed on square root scale.

^c P-value is based on difference of means on square root scale.

^b Slope and standard error based on square root of diastolic blood pressure versus log₂ (1987 dioxin + 1).

^b Slope and standard error based on square root of diastolic blood pressure versus log₂ (1987 dioxin + 1).

14.2.2.2.4 Diastolic Blood Pressure (Discrete)

All unadjusted and adjusted analyses of diastolic blood pressure in its dichotomous form were nonsignificant (Table 14-10(a–h): p>0.19 for each analysis).

Table 14-10. Analysis of Diastolic Blood Pressure (Discrete)

(a) MODEL 1: RANCH HANDS VS. COMPARISONS – UNADJUSTED					
Occupational Category	Group	n	Number (%) High	Est. Relative Risk (95% C.I.)	p-Value
All	Ranch Hand Comparison	859 1,232	45 (5.2) 61 (5.0)	1.06 (0.71,1.58)	0.769
Officer	Ranch Hand Comparison	334 484	20 (6.0) 22 (4.5)	1.34 (0.72,2.49)	0.360
Enlisted Flyer	Ranch Hand Comparison	149 186	8 (5.4) 8 (4.3)	1.26 (0.46,3.45)	0.649
Enlisted Groundcrew	Ranch Hand Comparison	376 562	17 (4.5) 31 (5.5)	0.81 (0.44,1.49)	0.499

(b) MODEL 1: RANCH HANDS VS. COMPARISONS – ADJUSTED					
Occurational Cotonsum	Adjusted Relative Risk	Volue			
Occupational Category	(95% C.I.)	p-Value			
All	1.02 (0.67,1.56)	0.916			
Officer	1.21 (0.62,2.35)	0.576			
Enlisted Flyer	1.18 (0.41,3.37)	0.760			
Enlisted Groundcrew	0.84 (0.44,1.59)	0.584			

(c) MODEL 2: RANCH HANDS – INITIAL DIOXIN – UNADJUSTED						
Initial I	Dioxin Category S	Summary Statistics	Analysis Results for Log ₂ (Initial Dioxin) ^a		
Initial Dioxin	n	Number (%) High	Estimated Relative Risk (95% C.I.) ^b	p-Value		
Low	155	7 (4.5)	1.04 (0.79,1.37)	0.793		
Medium	161	12 (7.5)				
High	160	10 (6.3)				

^a Adjusted for percent body fat at the time of the blood measurement of dioxin. ^b Relative risk for a twofold increase in initial dioxin.

(d) MODEL 2: RANCH HANDS – INITIAL DIOXIN – ADJUSTED				
	Analysis Results for Log ₂ (Initial Dioxin	n)		
	Adjusted Relative Risk			
n	(95% C.I.) ^a	p-Value		
457	1.15 (0.80,1.67)	0.446		

^a Relative risk for a twofold increase in initial dioxin.

Table 14-10. Analysis of Diastolic Blood Pressure (Discrete) (Continued)

(e) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY - UNADJUSTED

Dioxin Category	n	Number (%) High	Est. Relative Risk (95% C.I.) ^{ab}	p-Value
Comparison	1,195	59 (4.9)	(50 / 0 000)	p + mae
Background RH	376	15 (4.0)	0.85 (0.47,1.52)	0.576
Low RH	233	12 (5.2)	1.04 (0.55,1.96)	0.915
High RH	243	17 (7.0)	1.37 (0.78,2.41)	0.267
Low plus High RH	476	29 (6.1)	1.20 (0.75,1.90)	0.447

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: 1987 Dioxin ≤ 10 ppt.

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, 10 ppt < Initial Dioxin ≤ 94 ppt.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

(f) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY - ADJUSTED

	Adjusted Relative Risk			
Dioxin Category	n	" (95% C.I.) ^a	p-Value	
Comparison	1,155			
Background RH	360	0.78 (0.41,1.48)	0.449	
Low RH	221	0.91 (0.45,1.83)	0.792	
High RH	236	1.46 (0.80,2.68)	0.221	
Low plus High RH	457	1.16 (0.71,1.91)	0.551	

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: $1987 \text{ Dioxin} \le 10 \text{ ppt.}$

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, $10 \text{ ppt} < \text{Initial Dioxin} \le 94 \text{ ppt}$.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

(g) MODEL 4: RANCH HANDS – 1987 DIOXIN – UNADJUSTED

1987 Dioxin Category Summary Statistics			Analysis Results for Log ₂	(1987 Dioxin + 1)
1987 Dioxin	n	Number (%) High	Estimated Relative Risk (95% C.I.) ^a	p-Value
Low	284	14 (4.9)	1.14 (0.94,1.39)	0.198
Medium	281	9 (3.2)		
High	287	21 (7.3)		

^a Relative risk for a twofold increase in 1987 dioxin.

Note: Low = ≤ 7.9 ppt; Medium = > 7.9 - 19.6 ppt; High = > 19.6 ppt.

^b Adjusted for percent body fat at the time of the blood measurement of dioxin.

Table 14-10. Analysis of Diastolic Blood Pressure (Discrete) (Continued)

(h) MODEL 4: RANCH HANDS – 1987 DIOXIN – ADJUSTED					
Analysis Results for Log ₂ (1987 Dioxin + 1)					
Adjusted Relative Risk					
n	(95% C.I.) ^a	p-Value			
817	1.20 (0.89,1.61)	0.228			

^a Relative risk for a twofold increase in 1987 dioxin.

14.2.2.2.5 Heart Sounds

All Model 1 and Model 2 analyses of heart sounds were nonsignificant (Table 14-11(a-d): p>0.11 for each analysis).

Table 14-11. Analysis of Heart Sounds

(a) MODEL 1:	(a) MODEL 1: RANCH HANDS VS. COMPARISONS – UNADJUSTED					
Occupational Category	Group	n	Number (%) Abnormal	Est. Relative Risk (95% C.I.)	p-Value	
All	Ranch Hand Comparison	859 1,232	31 (3.6) 62 (5.0)	0.71 (0.45,1.10)	0.116	
Officer	Ranch Hand Comparison	334 484	11 (3.3) 26 (5.4)	0.60 (0.29,1.23)	0.164	
Enlisted Flyer	Ranch Hand Comparison	149 186	7 (4.7) 11 (5.9)	0.78 (0.30,2.08)	0.625	
Enlisted Groundcrew	Ranch Hand Comparison	376 562	13 (3.5) 25 (4.4)	0.77 (0.39,1.52)	0.452	

(b) MODEL 1: RANCH HANDS VS. COMPARISONS – ADJUSTED					
Occupational Category	Adjusted Relative Risk (95% C.I.)	p-Value			
All	0.71 (0.45,1.13)	0.139			
Officer	0.60 (0.28,1.29)	0.190			
Enlisted Flyer	0.65 (0.23,1.84)	0.419			
Enlisted Groundcrew	0.86 (0.42,1.74)	0.675			

(c) MODEL 2: RANCH HANDS – INITIAL DIOXIN – UNADJUSTED					
Initial I	Dioxin Category S	Summary Statistics	Analysis Results for Log ₂ (Initial Dioxin) ^a	
Initial Dioxin	n	Number (%) Abnormal	Estimated Relative Risk (95% C.I.) ^b	p-Value	
Low	155	6 (3.9)	1.01 (0.73,1.40)	0.958	
Medium	161	10 (6.2)			
High	160	6 (3.8)			

^a Adjusted for percent body fat at the time of the blood measurement of dioxin. ^b Relative risk for a twofold increase in initial dioxin.

Table 14-11. Analysis of Heart Sounds (Continued)

(d) MODEL 2: RANCH HANDS – INITIAL DIOXIN – ADJUSTED

(d) MODEL 2: RANCH HANDS – INITIAL DIOXIN – ADJUSTED					
Analysis Results for Log ₂ (Initial Dioxin)					
	Adjusted Relative Risk				
n	(95% C.I.) ^a	p-Value			
457	1.28 (0.83,1.98)	0.266			

^a Relative risk for a twofold increase in initial dioxin.

(e) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY - UNADJUSTED

Dioxin Category	n	Number (%) Abnormal	Est. Relative Risk (95% C.I.) ^{ab}	p-Value
Comparison	1,195	60 (5.0)		
Background RH	376	9 (2.4)	0.48 (0.24,0.99)	0.047
Low RH	233	10 (4.3)	0.84 (0.42,1.67)	0.622
High RH	243	12 (4.9)	0.94 (0.50,1.79)	0.857
Low plus High RH	476	22 (4.6)	0.89 (0.54,1.48)	0.656

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: $1987 \text{ Dioxin} \le 10 \text{ ppt.}$

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, $10 \text{ ppt} < \text{Initial Dioxin} \le 94 \text{ ppt}$.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

(f) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY - ADJUSTED

		Adjusted Relative Risk	
Dioxin Category	n	(95% C.I.) ^a	p-Value
Comparison	1,155		
Background RH	360	0.45 (0.21,0.97)	0.041
Low RH	221	0.80 (0.39,1.61)	0.528
High RH	236	1.05 (0.52,2.11)	0.901
Low plus High RH	457	0.92 (0.54,1.56)	0.750

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: $1987 \text{ Dioxin} \le 10 \text{ ppt.}$

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, 10 ppt < Initial Dioxin ≤ 94 ppt.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

^b Adjusted for percent body fat at the time of the blood measurement of dioxin.

Table 14-11. Analysis of Heart Sounds (Continued)

(g) MODEL 4: RANCH HANDS – 1987 DIOXIN – UNADJUSTED					
1987 Dioxin Category Summary Statistics			Analysis Results for Log ₂	(1987 Dioxin + 1)	
1987 Dioxin	n	Number (%) Abnormal	Estimated Relative Risk (95% C.I.) ^a	p-Value	
Low	284	8 (2.8)	1.16 (0.92,1.46)	0.220	

^a Relative risk for a twofold increase in 1987 dioxin.

281

287

Medium

High

Note: Low = ≤ 7.9 ppt; Medium = > 7.9 - 19.6 ppt; High = > 19.6 ppt.

9 (3.2)

14 (4.9)

(h) MODEL 4: RANCH HANDS – 1987 DIOXIN – ADJUSTED				
	Analysis Results for Log_2 (1987 Dioxin + 1)			
	Adjusted Relative Risk			
n	(95% C.I.) ^a	p-Value		
817	1.24 (0.89,1.73)	0.193		

^a Relative risk for a twofold increase in 1987 dioxin.

The unadjusted and adjusted Model 3 analyses each showed a significant difference between Ranch Hands in the background dioxin category and Comparisons (Table 14-11(e,f): Est. RR=0.48, p=0.047; Adj. RR=0.45, p=0.041, respectively). The percentage of participants with abnormal heart sounds was lower for Ranch Hands in the background dioxin category (2.4%) than for Comparisons (5.0%).

Model 4 unadjusted and adjusted analyses showed no significant association between heart sounds and 1987 dioxin (Table 14-11(g,h): p>0.19 for each analysis).

14.2.2.2.6 Overall Electrocardiograph

The unadjusted and adjusted Model 1 analyses of overall ECG showed no overall group difference between Ranch Hands and Comparisons (Table 14-12(a,b): p>0.68 for each contrast). Stratifying by occupation revealed a marginally significant group difference within the enlisted groundcrew stratum for both the unadjusted and adjusted analyses (Table 14-12(a,b): Est. RR=0.77, p=0.096; Adj. RR=0.76, p=0.095, respectively). The percentage of enlisted groundcrew with abnormal overall ECG results was lower for Ranch Hands (23.4%) than for Comparisons (28.3%).

Both the unadjusted and adjusted Model 2 analyses of overall ECG were nonsignificant (Table 14-12(c,d): p>0.17 for each analysis).

The unadjusted Model 3 analyses of overall ECG did not show any of the Ranch Hand categories to be significantly different from the Comparison group (Table 14-12(e): p>0.60 for each contrast). After adjusting for covariates, a marginally significant difference between Ranch Hands in the low dioxin category and Comparisons was revealed (Table 14-12(f): Adj. RR=0.73, p=0.063). The percentage of abnormal overall ECG results was lower for Ranch Hands (30.9%) than for Comparisons (31.2%). Both the unadjusted and adjusted Model 4 analyses were nonsignificant (Table 14-12(g,h): p>0.39 for each analysis).

Table 14-12. Analysis of Overall Electrocardiograph (ECG)

(a) MODEL 1: RANCH HANDS VS. COMPARISONS - UNADJUSTED

Occupational Category	Group	n	Number (%) Abnormal	Est. Relative Risk (95% C.I.)	p-Value
All	Ranch Hand Comparison	859 1,232	268 (31.2) 384 (31.2)	1.00 (0.83,1.21)	0.988
Officer	Ranch Hand Comparison	334 484	120 (35.9) 163 (33.7)	1.10 (0.82,1.48)	0.506
Enlisted Flyer	Ranch Hand Comparison	149 186	60 (40.3) 62 (33.3)	1.35 (0.86,2.11)	0.190
Enlisted Groundcrew	Ranch Hand Comparison	376 562	88 (23.4) 159 (28.3)	0.77 (0.57,1.05)	0.096

(b) MODEL 1: RANCH HANDS VS. COMPARISONS - ADJUSTED

	Adjusted Relative Risk	
Occupational Category	(95% C.I.)	p-Value
All	0.96 (0.78,1.18)	0.688
Officer	1.07 (0.79,1.47)	0.655
Enlisted Flyer	1.24 (0.76,2.00)	0.389
Enlisted Groundcrew	0.76 (0.55,1.05)	0.095

(c) MODEL 2: RANCH HANDS - INITIAL DIOXIN - UNADJUSTED

Initial Dioxin Category Summary Statistics			Analysis Results for Log ₂ (Initial Dioxin) ^a
Initial Dioxin	n	Number (%) Abnormal	Estimated Relative Risk (95% C.I.) ^b	p-Value
Low	155	51 (32.9)	0.90 (0.77,1.05)	0.171
Medium	161	47 (29.2)		
High	160	48 (30.0)		

^a Adjusted for percent body fat at the time of the blood measurement of dioxin. ^b Relative risk for a twofold increase in initial dioxin.

Note: Low = 27-63 ppt; Medium = >63-152 ppt; High = >152 ppt.

(d) MODEL 2: RANCH HANDS – INITIAL DIOXIN – ADJUSTED

	Analysis Results for Log ₂ (Initial Diox	in)
	Adjusted Relative Risk	
n	(95% C.I.) ^a	p-Value
457	1.14 (0.93,1.39)	0.200

^a Relative risk for a twofold increase in initial dioxin.

Table 14-12. Analysis of Overall Electrocardiograph (ECG) (Continued)

(e) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY - UNADJUSTED

		Number (%)	Est. Relative Risk	
Dioxin Category	n	Abnormal	(95% C.I.) ^{ab}	p-Value
Comparison	1,195	373 (31.2)		
Background RH	376	118 (31.4)	1.06 (0.82,1.36)	0.659
Low RH	233	72 (30.9)	0.98 (0.72,1.33)	0.883
High RH	243	74 (30.5)	0.92 (0.68,1.25)	0.602
Low plus High RH	476	146 (30.7)	0.95 (0.75,1.20)	0.659

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: 1987 Dioxin ≤ 10 ppt.

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, 10 ppt < Initial Dioxin ≤ 94 ppt.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

(f) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY - ADJUSTED

		Adjusted Relative Risk	
Dioxin Category	n	(95% C.I.) ^a	p-Value
Comparison	1,155		
Background RH	360	1.00 (0.76,1.32)	0.980
Low RH	221	0.73 (0.52,1.02)	0.063
High RH	236	1.10 (0.78,1.54)	0.578
Low plus High RH	457	0.90 (0.70,1.16)	0.423

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: $1987 \text{ Dioxin} \le 10 \text{ ppt.}$

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, $10 \text{ ppt} < \text{Initial Dioxin} \le 94 \text{ ppt}$.

 $High \; (Ranch \; Hand) \colon \; 1987 \; Dioxin > 10 \; ppt, \; Initial \; Dioxin > 94 \; ppt.$

(g) MODEL 4: RANCH HANDS - 1987 DIOXIN - UNADJUSTED

1987 Dioxin Category Summary Statistics			Analysis Results for Log ₂	(1987 Dioxin + 1)
1987 Dioxin	n	Number (%) Abnormal	Estimated Relative Risk (95% C.I.) ^a	p-Value
Low	284	84 (29.6)	0.96 (0.87,1.06)	0.391
Medium	281	93 (33.1)		
High	287	87 (30.3)		

^a Relative risk for a twofold increase in 1987 dioxin.

Note: Low = ≤ 7.9 ppt; Medium = > 7.9 - 19.6 ppt; High = > 19.6 ppt.

^b Adjusted for percent body fat at the time of the blood measurement of dioxin.

Table 14-12. Analysis of Overall Electrocardiograph (ECG) (Continued)

(h) MODEL 4: RANCH HANDS – 1987 DIOXIN – ADJUSTED					
	Analysis Results for Log ₂ (1987 Dioxin + 1)				
	Adjusted Relative Risk				
n	(95% C.I.) ^a	p-Value			
817	1.02 (0.89,1.17)	0.753			

^a Relative risk for a twofold increase in 1987 dioxin.

14.2.2.2.7 Right Bundle Branch Block

All unadjusted and adjusted analysis results of right bundle branch block were nonsignificant (Table 14-13(a–h): p>0.27 for each analysis).

Table 14-13. Analysis of Right Bundle Branch Block

(a) MODEL 1: RANCH HANDS VS. COMPARISONS – UNADJUSTED					
Occupational Category	Group	n	Number (%) Yes	Est. Relative Risk (95% C.I.)	p-Value
All	Ranch Hand Comparison	859 1,232	21 (2.4) 33 (2.7)	0.91 (0.52,1.58)	0.739
Officer	Ranch Hand Comparison	334 484	8 (2.4) 13 (2.7)	0.89 (0.36,2.17)	0.796
Enlisted Flyer	Ranch Hand Comparison	149 186	8 (5.4) 7 (3.8)	1.45 (0.51,4.10)	0.482
Enlisted Groundcrew	Ranch Hand Comparison	376 562	5 (1.3) 13 (2.3)	0.57 (0.20,1.61)	0.288

(b) MODEL 1: RANCH HANDS VS. COMPARISONS – ADJUSTED					
Adjusted Relative Risk Occupational Category (95% C.I.) p-Value					
All	0.88 (0.49,1.56)	0.650			
Officer	0.89 (0.36,2.22)	0.807			
Enlisted Flyer	1.47 (0.49,4.44)	0.493			
Enlisted Groundcrew	0.55 (0.19,1.59)	0.271			

(c) MODEL 2: RANCH HANDS – INITIAL DIOXIN – UNADJUSTED					
Initial Dioxin Category Summary Statistics Analysis Results for Log ₂ (Initial Dioxin) ^a					
Initial Dioxin	n	Number (%) Yes	Estimated Relative Risk (95% C.I.) ^b	p-Value	
Low	155	5 (3.2)	0.93 (0.59,1.46)	0.747	
Medium	161	4 (2.5)			
High	160	3 (1.9)			

^a Adjusted for percent body fat at the time of the blood measurement of dioxin.

^b Relative risk for a twofold increase in initial dioxin.

Table 14-13. Analysis of Right Bundle Branch Block (Continued)

(d) MODEL 2: RANCH HANDS – INITIAL DIOXIN – ADJUSTED				
	Analysis Results for Log ₂ (Initial Diox	cin)		
	Adjusted Relative Risk			
n	(95% C.I.) ^a	p-Value		
462	1.12 (0.62.2.04)	0.707		

^a Relative risk for a twofold increase in initial dioxin.

Note: Results are not adjusted for race and family history of heart disease before age 45 because of the sparse number of Ranch Hands with a right bundle branch block.

(e) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY – UNADJUSTED

		Number (%)	Est. Relative Risk	
Dioxin Category	n	Yes	(95% C.I.) ^{ab}	p-Value
Comparison	1,195	31 (2.6)		
Background RH	376	9 (2.4)	0.93 (0.44,1.98)	0.852
Low RH	233	5 (2.1)	0.82 (0.32,2.14)	0.688
High RH	243	7 (2.9)	1.10 (0.48,2.54)	0.818
Low plus High RH	476	12 (2.5)	0.96 (0.48,1.89)	0.895

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: 1987 Dioxin ≤ 10 ppt.

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, 10 ppt < Initial Dioxin ≤ 94 ppt.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

(f) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY - ADJUSTED

		Adjusted Relative Risk	
Dioxin Category	n	(95% C.I.) ^a	p-Value
Comparison	1,155		
Background RH	360	1.04 (0.47,2.29)	0.920
Low RH	221	0.55 (0.19,1.60)	0.273
High RH	236	1.19 (0.49,2.88)	0.704
Low plus High RH	457	0.82 (0.39,1.71)	0.594

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: $1987 \text{ Dioxin} \le 10 \text{ ppt.}$

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, 10 ppt < Initial Dioxin ≤ 94 ppt.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

^b Adjusted for percent body fat at the time of the blood measurement of dioxin.

Table 14-13. Analysis of Right Bundle Branch Block (Continued)

(g) MODEL 4: RANCH HANDS – 1987 DIOXIN – UNADJUSTED				
1987 Dioxin Category Summary Statistics			Analysis Results for Log ₂	(1987 Dioxin + 1)
1987 Dioxin	n	Number (%) Yes	Estimated Relative Risk (95% C.I.) ^a	p-Value
Low	284	6 (2.1)	1.03 (0.77,1.38)	0.845
Medium	281	8 (2.8)		
High	287	7 (2.4)		

^a Relative risk for a twofold increase in 1987 dioxin.

Note: Low = ≤ 7.9 ppt; Medium = > 7.9 - 19.6 ppt; High = > 19.6 ppt.

(h) MODEL 4: RANCH HANDS – 1987 DIOXIN – ADJUSTED				
	Analysis Results for Log ₂ (1987 Dioxin + 1)			
	Adjusted Relative Risk			
n	(95% C.I.) ^a	p-Value		
817	1.02 (0.69,1.50)	0.922		

^a Relative risk for a twofold increase in 1987 dioxin.

Note: Results are not adjusted for race because of the sparse number of Ranch Hands with a right bundle branch block.

14.2.2.2.8 Left Bundle Branch Block

The unadjusted and adjusted Model 1 analyses of left bundle branch block were nonsignificant (Table 14-14(a,b): p≥0.15 for each contrast).

Table 14-14. Analysis of Left Bundle Branch Block

(a) MODEL 1: RANCH HANDS VS. COMPARISONS – UNADJUSTED					
Occupational Category	Group	n	Number (%) Yes	Est. Relative Risk (95% C.I.)	p-Value
All	Ranch Hand Comparison	859 1,232	5 (0.6) 12 (1.0)	0.60 (0.21,1.70)	0.317
Officer	Ranch Hand Comparison	334 484	2 (0.6) 6 (1.2)	0.48 (0.10,2.39)	0.370
Enlisted Flyer	Ranch Hand Comparison	149 186	1 (0.7) 0 (0.0)		0.911 ^a
Enlisted Groundcrew	Ranch Hand Comparison	376 562	2 (0.5) 6 (1.1)	0.50 (0.10,2.47)	0.391

^a P-value determined using a chi-square test with continuity correction because of the sparse number of participants with a left bundle branch block.

^{--:} Results not presented because of the sparse number of participants with a left bundle branch block.

Table 14-14. Analysis of Left Bundle Branch Block (Continued)

(b) MODEL 1: RANCH HANDS VS. COMPARISONS - ADJUSTED

(6) 110222 11 121 (51 121 122) 61 (61 121 122 61 61 122 61 122 61 122 61 122 61 122 61 122 61 122 61 122 61 122 61 122 61 61 122 61 61 61 61 61 61 61 61 61 61 61 61 61						
Adjusted Relative Risk						
Occupational Category	(95% C.I.)	p-Value				
All	0.47 (0.15,1.50)	0.182				
Officer	0.21 (0.02,1.76)	0.150				
Enlisted Flyer						
Enlisted Groundcrew	0.56 (0.11,2.83)	0.479				

^{--:} Results not presented because of the sparse number of participants with a left bundle branch block.

Note: Results are not adjusted for race because of the sparse number of participants with a left bundle branch block.

(c) MODEL 2: RANCH HANDS – INITIAL DIOXIN – UNADJUSTED

Initial Dioxin Category Summary Statistics		Analysis Results for Log ₂ ((nitial Dioxin) ^a	
Initial Dioxin	n	Number (%) Yes	Estimated Relative Risk (95% C.I.) ^b	p-Value
Low	155	1 (0.6)	0.21 (0.01,6.22)	0.213
Medium	161	0(0.0)		
High	160	0 (0.0)		

^a Adjusted for percent body fat at the time of the blood measurement of dioxin.

Note: Low = 27-63 ppt; Medium = >63-152 ppt; High = >152 ppt.

(d) MODEL 2: RANCH HANDS – INITIAL DIOXIN – ADJUSTED

	Analysis Results for Log ₂ (Initial Dioxin	
n	Adjusted Relative Risk (95% C.I.)	p-Value

^{--:} Results not presented because of the sparse number of Ranch Hands with a left bundle branch block.

(e) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY – UNADJUSTED

		Number (%)	Est. Relative Risk	
Dioxin Category	n	Yes	(95% C.I.) ^{ab}	p-Value
Comparison	1,195	12 (1.0)		
Background RH	376	4 (1.1)	1.17 (0.37,3.68)	0.792
Low RH	233	1 (0.4)	0.42 (0.05,3.23)	0.403
High RH	243	0 (0.0)		0.237^{c}
Low plus High RH	476	1 (0.2)		0.174^{c}

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: $1987 \text{ Dioxin} \le 10 \text{ ppt.}$

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, $10 \text{ ppt} < \text{Initial Dioxin} \le 94 \text{ ppt}$.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

^b Relative risk for a twofold increase in initial dioxin.

^b Adjusted for percent body fat at the time of the blood measurement of dioxin.

^c P-value determined using a chi-square test with continuity correction because of the sparse number of participants with left bundle branch block.

^{--:} Results not presented because of the sparse number of participants with a left bundle branch block.

Table 14-14. Analysis of Left Bundle Branch Block (Continued)

(f) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY – ADJUSTED				
Dioxin Category	n	Adjusted Relative Risk (95% C.I.) ^a	p-Value	
Comparison	1,155			
Background RH	360	0.87 (0.23,3.33)	0.838	
Low RH	221	0.37 (0.05,2.91)	0.341	
High RH	236			
Low plus High RH	457			

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: $1987 \text{ Dioxin} \le 10 \text{ ppt.}$

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, 10 ppt < Initial Dioxin ≤ 94 ppt.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

Results are not adjusted for race because of the sparse number of participants with a left bundle branch block.

(g) MODEL 4: RANCH HANDS – 1987 DIOXIN – UNADJUSTED					
1987 Dioxin Category Summary Statistics			Analysis Results for Log ₂	(1987 Dioxin + 1)	
1987		Number (%)	Estimated Relative Risk	<u></u> .	
Dioxin	n	Yes	(95% C.I.) ^a	p-Value	
Low	284	1 (0.4)	0.69 (0.35,1.36)	0.271	
Medium	281	4 (1.4)			
High	287	0 (0.0)			

^a Relative risk for a twofold increase in 1987 dioxin.

Note: Low = ≤ 7.9 ppt; Medium = > 7.9 - 19.6 ppt; High = > 19.6 ppt.

(h) MODEL 4: RANCH HANDS – 1987 DIOXIN – ADJUSTED				
	Analysis Results for Log_2 (1987 Dioxin + 1)			
Adjusted Relative Risk				
n	(95% C.I.) ^a	p-Value		
823	0.56 (0.23,1.39)	0.199		

^a Relative risk for a twofold increase in 1987 dioxin.

Note: Results are not adjusted for race and diabetic class because of the sparse number of participants with a left bundle branch block.

The unadjusted Model 2 analysis showed no significant association between left bundle branch block and initial dioxin (Table 14-14(c): p=0.213). Because of a sparse number of Ranch Hands with a left bundle branch block, the adjusted Model 2 analysis was not performed.

All unadjusted and adjusted Model 3 and 4 analyses were nonsignificant (Table 14-14(e-h): p>0.17 for each analysis).

^{--:} Results not presented because of the sparse number of participants with a left bundle branch block.

14.2.2.2.9 Non-Specific ST- and T-Wave Changes

All unadjusted and adjusted analyses of non-specific ST- and T-wave changes were nonsignificant (Table 14-15(a-h): p≥0.18 for each analysis).

Table 14-15. Analysis of Non-Specific ST- and T-Wave Changes

Occupational Category	Group	n	Number (%) Yes	Est. Relative Risk (95% C.I.)	p-Value
All	Ranch Hand Comparison	859 1,232	160 (18.6) 222 (18.0)	1.04 (0.83,1.30)	0.724
Officer	Ranch Hand Comparison	334 484	70 (21.0) 95 (19.6)	1.09 (0.77,1.53)	0.641
Enlisted Flyer	Ranch Hand Comparison	149 186	33 (22.1) 34 (18.3)	1.27 (0.74,2.17)	0.380
Enlisted Groundcrew	Ranch Hand Comparison	376 562	57 (15.2) 93 (16.5)	0.90 (0.63,1.29)	0.570

(b) MODEL 1: RANCH HANDS VS. COMPARISONS – ADJUSTED					
Occupational Category	Adjusted Relative Risk (95% C.I.)	p-Value			
All	1.00 (0.79,1.27)	0.984			
Officer	1.03 (0.71,1.48)	0.882			
Enlisted Flyer	1.22 (0.69,2.14)	0.495			
Enlisted Groundcrew	0.88 (0.60,1.29)	0.517			

(c) MODEL 2: RANCH HANDS – INITIAL DIOXIN – UNADJUSTED						
Initial I	Analysis Results for Log ₂ (Initial Dioxin) ^a				
Initial Dioxin	n	Number (%) Yes	Estimated Relative Risk (95% C.I.) ^b	p-Value		
Low	155	32 (20.6)	0.91 (0.76,1.08)	0.280		
Medium	161	34 (21.1)				
High	160	31 (19.4)				

^a Adjusted for percent body fat at the time of the blood measurement of dioxin. ^b Relative risk for a twofold increase in initial dioxin.

(d) MODEL 2: RANCH l	HANDS – INITIAL DIOXIN – ADJUSTED	
	Analysis Results for Log ₂ (Initial Dioxi	n)
	Adjusted Relative Risk	
n	(95% C.I.) ^a	p-Value
457	1.15 (0.91,1.44)	0.237

^a Relative risk for a twofold increase in initial dioxin.

Table 14-15. Analysis of Non-Specific ST- and T-Wave Changes (Continued)

(e) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY - UNADJUSTED

		Number (%)	Est. Relative Risk	
Dioxin Category	n	Yes	(95% C.I.) ^{ab}	p-Value
Comparison	1,195	218 (18.2)		
Background RH	376	59 (15.7)	0.91 (0.66,1.25)	0.545
Low RH	233	47 (20.2)	1.12 (0.78,1.59)	0.537
High RH	243	50 (20.6)	1.08 (0.76,1.52)	0.677
Low plus High RH	476	97 (20.4)	1.10 (0.84,1.44)	0.502

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: 1987 Dioxin ≤ 10 ppt.

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, 10 ppt < Initial Dioxin ≤ 94 ppt.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

(f) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY - ADJUSTED

		Adjusted Relative Risk	
Dioxin Category	n	(95% C.I.) ^a	p-Value
Comparison	1,155		
Background RH	360	0.82 (0.58,1.15)	0.242
Low RH	221	0.91 (0.62,1.32)	0.614
High RH	236	1.26 (0.86,1.84)	0.238
Low plus High RH	457	1.07 (0.80,1.43)	0.628

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: $1987 \text{ Dioxin} \le 10 \text{ ppt.}$

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, $10 \text{ ppt} < \text{Initial Dioxin} \le 94 \text{ ppt}$.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

(g) MODEL 4: RANCH HANDS – 1987 DIOXIN – UNADJUSTED

1987 Dioxin Category Summary Statistics		Analysis Results for Log ₂	(1987 Dioxin + 1)	
1987 Dioxin	n	Number (%) Yes	Estimated Relative Risk (95% C.I.) ^a	p-Value
Low	284	43 (15.1)	1.06 (0.94,1.19)	0.361
Medium	281	52 (18.5)		
High	287	61 (21.3)		

^a Relative risk for a twofold increase in 1987 dioxin.

Note: Low = ≤ 7.9 ppt; Medium = > 7.9 - 19.6 ppt; High = > 19.6 ppt.

^b Adjusted for percent body fat at the time of the blood measurement of dioxin.

Table 14-15. Analysis of Non-Specific ST- and T-Wave Changes (Continued)

(h) MODEL 4: RANCH HANDS – 1987 DIOXIN – ADJUSTED				
	Analysis Results for Log ₂ (1987 Dioxin + 1)			
Adjusted Relative Risk				
n	(95% C.I.) ^a	p-Value		
817	1.12 (0.95,1.32)	0.180		

^a Relative risk for a twofold increase in 1987 dioxin.

14.2.2.2.10 Bradycardia

The Model 1 and 2 analyses of bradycardia did not show a significant association with dioxin in either the unadjusted or adjusted analysis (Table 14-16(a–d): p≥0.12 for each analysis).

Table 14-16. Analysis of Bradycardia

(a) MODEL 1:	(a) MODEL 1: RANCH HANDS VS. COMPARISONS – UNADJUSTED					
Occupational Category	Group	n	Number (%) Yes	Est. Relative Risk (95% C.I.)	p-Value	
All	Ranch Hand Comparison	859 1,232	24 (2.8) 49 (4.0)	0.69 (0.42,1.14)	0.142	
Officer	Ranch Hand Comparison	334 484	15 (4.5) 31 (6.4)	0.69 (0.36,1.29)	0.245	
Enlisted Flyer	Ranch Hand Comparison	149 186	5 (3.4) 5 (2.7)	1.26 (0.36,4.43)	0.722	
Enlisted Groundcrew	Ranch Hand Comparison	376 562	4 (1.1) 13 (2.3)	0.45 (0.15,1.40)	0.170	

(b) MODEL 1: RANCH HANDS VS. COMPARISONS – ADJUSTED					
Adjusted Relative Risk Occupational Category (95% C.I.) p-Value					
All	0.69 (0.41,1.16)	0.151			
Officer	0.74 (0.38,1.42)	0.360			
Enlisted Flyer	1.14 (0.32,4.09)	0.846			
Enlisted Groundcrew	0.36 (0.10,1.30)	0.120			

(c) MODEL 2: RANCH HANDS – INITIAL DIOXIN – UNADJUSTED					
Initial Dioxin Category Summary Statistics Analysis Results for Log ₂ (Initial Dioxin) ^a					
Initial		Number (%)			
Dioxin	n	Yes	(95% C.I.) ^b	p-Value	
Low	155	4 (2.6)	0.86 (0.44,1.65)	0.631	
Medium	161	2 (1.2)			
High	160	1 (0.6)			

^a Adjusted for percent body fat at the time of the blood measurement of dioxin. ^b Relative risk for a twofold increase in initial dioxin.

Table 14-16. Analysis of Bradycardia (Continued)

(d) MODEL 2: RANCH HANDS – INITIAL DIOXIN – ADJUSTED					
	Analysis Results for Log ₂ (Initial Diox	in)			
	Adjusted Relative Risk				
n	(95% C.I.) ^a	p-Value			
466	0.98 (0.44.2.22)	0.971			

^a Relative risk for a twofold increase in initial dioxin.

Note: Results are not adjusted for race, diabetic class, and family history of heart disease before age 45 because of the sparse number of Ranch Hands with bradycardia.

(e) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY - UNADJUSTED

		Number (%)	Est. Relative Risk	
Dioxin Category	n	Yes	(95% C.I.) ^{ab}	p-Value
Comparison	1,195	47 (3.9)		
Background RH	376	16 (4.3)	0.95 (0.53,1.71)	0.867
Low RH	233	5 (2.1)	0.55 (0.21,1.39)	0.204
High RH	243	2 (0.8)	0.23 (0.05, 0.95)	0.042
Low plus High RH	476	7 (1.5)	0.35 (0.14,0.85)	0.020

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: 1987 Dioxin ≤ 10 ppt.

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, 10 ppt < Initial Dioxin ≤ 94 ppt.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

(f) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY - ADJUSTED

		Adjusted Relative Risk	
Dioxin Category	n	(95% C.I.) ^a	p-Value
Comparison	1,155		
Background RH	360	0.81 (0.44,1.49)	0.497
Low RH	221	0.49 (0.17,1.40)	0.183
High RH	236	0.35 (0.08,1.50)	0.156
Low plus High RH	457	0.41 (0.16,1.05)	0.062

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: $1987 \text{ Dioxin} \le 10 \text{ ppt.}$

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, 10 ppt < Initial Dioxin ≤ 94 ppt.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

^b Adjusted for percent body fat at the time of the blood measurement of dioxin.

Table 14-16. Analysis of Bradycardia (Continued)

(g) MODEL 4:	(g) MODEL 4: RANCH HANDS – 1987 DIOXIN – UNADJUSTED						
1987 Dioxir	1987 Dioxin Category Summary Statistics Analysis Results for Log ₂ (1987 Dioxin + 1)						
1987 Dioxin	n	Number (%) Yes	Estimated Relative Risk (95% C.I.) ^a	p-Value			
Low	284	11 (3.9)	0.77 (0.56,1.05)	0.084			
Medium	281	9 (3.2)					
High	287	3 (1.0)					

^a Relative risk for a twofold increase in 1987 dioxin.

Note: Low = ≤ 7.9 ppt; Medium = > 7.9 - 19.6 ppt; High = > 19.6 ppt.

(h) MODEL 4: RANCH HANDS – 1987 DIOXIN – ADJUSTED						
	Analysis Results for Log ₂ (1987 Dioxin + 1)					
	Adjusted Relative Risk					
n	(95% C.I.) ^a	p-Value				
828	0.98 (0.65,1.49)	0.932				

^a Relative risk for a twofold increase in 1987 dioxin.

Note: Results are not adjusted for family history of heart disease before age 45 because of the sparse number of Ranch Hands with bradycardia.

The unadjusted Model 3 analysis of bradycardia revealed two significant contrasts: Ranch Hands in the high dioxin category versus Comparisons and Ranch Hands in the low plus high dioxin category versus Comparisons (Table 14-16(e): Est. RR=0.23, p=0.042; Est. RR=0.35, p=0.020, respectively). The percentage of participants with bradycardia was higher for Comparisons (3.9%) than for Ranch Hands in the high dioxin category (0.8%) or Ranch Hands in the low plus high dioxin category (1.5%). After covariate adjustment, there was a marginally significant difference between Ranch Hands in the low plus high dioxin category and Comparisons (Table 14-16(f): Adj. RR=0.41, p=0.062).

The unadjusted Model 4 analysis of bradycardia revealed a marginally significant inverse association between bradycardia and 1987 dioxin (Table 14-16(g): Est. RR=0.77, p=0.084). The percentages of participants with bradycardia in the low, medium, and high 1987 dioxin categories were 3.9, 3.2, and 1.0, respectively. After covariate adjustment, the results became nonsignificant (Table 14-16(h): p=0.932).

14.2.2.2.11 Tachycardia

The unadjusted and adjusted Model 1 analyses of tachycardia were nonsignificant (Table 14-17(a,b): p>0.12 for each contrast).

Table 14-17. Analysis of Tachycardia

() 3.50 DET 4	DANGER TEANING TIG	COLEDIDAGONG	TINI A TO THE COMPANY
(a) MODEL 1:	: RANCH HANDS VS.	COMPARISONS -	- UNADJUSTED

Occupational Category	Group	n	Number (%) Yes	Est. Relative Risk (95% C.I.)	p-Value
All	Ranch Hand Comparison	859 1,232	6 (0.7) 4 (0.3)	2.16 (0.61,7.68)	0.228
Officer	Ranch Hand Comparison	334 484	1 (0.3) 1 (0.2)	1.45 (0.09,23.27)	0.793
Enlisted Flyer	Ranch Hand Comparison	149 186	3 (2.0) 0 (0.0)		0.174 ^a
Enlisted Groundcrew	Ranch Hand Comparison	376 562	2 (0.5) 3 (0.5)	1.00 (0.17,5.99)	0.997

^a P-value determined using a chi-square test with continuity correction because of the sparse number of participants with tachycardia.

^{--:} Results not presented because of the sparse number of participants with tachycardia.

(b) MODEL 1: RANCH HANDS VS. COMPARISONS – ADJUSTED				
	Adjusted Relative Risk			
Occupational Ca	ntegory (95% C.I.)	p-Value		

 Occupational Category
 (95% C.I.)
 p-Value

 All
 2.94 (0.69,12.51)
 0.129

 Officer
 - -

 Enlisted Flyer
 - -

 Enlisted Groundcrew
 1.54 (0.19,12.63)
 0.685

Note: Results are not adjusted for family history of heart disease because of the sparse number of participants with tachycardia.

(c) MODEL 2: RANCH HANDS – INITIAL DIOXIN – UNADJUSTED

Initial Dioxin Category Summary Statistics		Analysis Results for Log ₂ (Initial Dioxin) ^a	
Initial Dioxin	n	Number (%) Yes	Estimated Relative Risk (95% C.I.) ^b	p-Value
Low	155	0 (0.0)	1.38 (0.72,2.68)	0.340
Medium	161	1 (0.6)		
High	160	3 (1.9)		

^a Adjusted for percent body fat at the time of the blood measurement of dioxin.

^{--:} Results not presented because of the sparse number of participants with tachycardia.

^b Relative risk for a twofold increase in initial dioxin.

Table 14-17. Analysis of Tachycardia (Continued)

(d) MODEL 2: RANCH HANDS – INITIAL DIOXIN – ADJUSTED Analysis Results for Log₂ (Initial Dioxin) Adjusted Relative Risk n (95% C.I.) p-Value

(e) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY - UNADJUSTED Number (%) Est. Relative Risk (95% C.I.)ab **Dioxin Category** Yes p-Value n Comparison 1.195 3(0.3)Background RH 1 (0.3) 376 1.33 (0.14,13.00) 0.806 Low RH 0.999^{c} 233 0(0.0)High RH 243 4 (1.6) 5.30 (1.15,24.53) 0.033 Low plus High RH 476 4(0.8) 0.206°

Note: RH = Ranch Hand.

Comparison: $1987 \text{ Dioxin} \le 10 \text{ ppt.}$

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, 10 ppt < Initial Dioxin ≤ 94 ppt.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

(f) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY - ADJUSTED

		Adjusted Relative Risk	
Dioxin Category	n	(95% C.I.) ^a	p-Value
Comparison	1,155		
Background RH	360	2.01 (0.16,24.61)	0.585
Low RH	221		
High RH	236	8.10 (1.19,55.01)	0.032
Low plus High RH	457		

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: $1987 \text{ Dioxin} \le 10 \text{ ppt.}$

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, $10 \text{ ppt} < \text{Initial Dioxin} \le 94 \text{ ppt}$.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

Results are not adjusted for family history of heart disease because of the sparse number of participants with tachycardia.

^{--:} Results not presented because of the sparse number of Ranch Hands with tachycardia.

^a Relative risk and confidence interval relative to Comparisons.

^b Adjusted for percent body fat at the time of the blood measurement of dioxin.

^c P-value determined using a chi-square test with continuity correction because of the sparse number of participants with tachycardia.

^{--:} Results not presented because of the sparse number of participants with tachycardia.

^{--:} Results not presented because of the sparse number of participants with tachycardia.

Table 14-17. Analysis of Tachycardia (Continued)

(g) MODEL 4:	(g) MODEL 4: RANCH HANDS – 1987 DIOXIN – UNADJUSTED						
1987 Dioxin Category Summary Statistics			Analysis Results for Log ₂	(1987 Dioxin + 1)			
1987		Number (%)	Estimated Relative Risk				
Dioxin	n	Yes	(95% C.I.) ^a	p-Value			
Low	284	1 (0.4)	1.56 (0.92,2.63)	0.111			
Medium	281	0 (0.0)					
High	287	4 (1.4)					

^a Relative risk for a twofold increase in 1987 dioxin.

Note: Low = ≤ 7.9 ppt; Medium = > 7.9 - 19.6 ppt; High = > 19.6 ppt.

(h) MODEL 4: RANCH HANDS	– 1987 DIOXIN – ADJUSTED	
	Analysis Results for Log ₂ (1987 Dioxin + 1)	
	Adjusted Relative Risk	
n	(95% C.I.) ^a	p-Value
825	1.55 (0.85,2.84)	0.165

^a Relative risk for a twofold increase in 1987 dioxin.

Note: Results are not adjusted for occupation, current alcohol use, personality type, family history of heart disease, and diabetic class because of the sparse number of participants with tachycardia.

The unadjusted Model 2 analysis showed no significant association between tachycardia and initial dioxin (Table 14-17(c): p=0.340). Because of a sparse number of Ranch Hands with tachycardia, the adjusted Model 2 analysis was not performed.

The unadjusted and adjusted Model 3 analyses each showed a significant difference between Ranch Hands in the high dioxin category and Comparisons (Table 14-17(e,f): Est. RR=5.30, p=0.033; Adj. RR=8.10, p=0.032, respectively). The percentage of participants with tachycardia for Ranch Hands in the high dioxin categories was 1.6 versus 0.3 percent for Comparisons.

The unadjusted and adjusted Model 4 analyses were nonsignificant (Table 14-17(g,h): p>0.11 for each analysis).

14.2.2.2.12 Arrhythmia

All unadjusted and adjusted analyses of arrhythmia were nonsignificant (Table 14-18(a–h): p>0.11 for each analysis).

Table 14-18. Analysis of Arrhythmia

(a) MODEL 1:	RANCH HANDS VS.	COMPARISONS -	UNADJUSTED

Occupational Category	Group	n	Number (%) Yes	Est. Relative Risk (95% C.I.)	p-Value
All	Ranch Hand Comparison	859 1,232	51 (5.9) 68 (5.5)	1.08 (0.74,1.57)	0.686
Officer	Ranch Hand Comparison	334 484	25 (7.5) 25 (5.2)	1.49 (0.84,2.63)	0.176
Enlisted Flyer	Ranch Hand Comparison	149 186	13 (8.7) 12 (6.5)	1.39 (0.61,3.13)	0.433
Enlisted Groundcrew	Ranch Hand Comparison	376 562	13 (3.5) 31 (5.5)	0.61 (0.32,1.19)	0.147

(b) MODEL 1: RANCH HANDS VS. COMPARISONS – ADJUSTED

Occupational Category	Adjusted Relative Risk (95% C.I.)	p-Value
All	1.02 (0.69,1.52)	0.913
Officer	1.39 (0.75,2.55)	0.296
Enlisted Flyer	1.26 (0.54,2.97)	0.591
Enlisted Groundcrew	0.62 (0.31,1.25)	0.180

(c) MODEL 2: RANCH HANDS – INITIAL DIOXIN – UNADJUSTED

Initial Dioxin Category Summary Statistics		Analysis Results for Log ₂ (Initial Dioxin) ^a		
Initial Dioxin	n	Number (%) Yes	Estimated Relative Risk (95% C.I.) ^b	p-Value
Low	155	13 (8.4)	0.81 (0.60,1.10)	0.158
Medium	161	11 (6.8)		
High	160	8 (5.0)		

^a Adjusted for percent body fat at the time of the blood measurement of dioxin. ^b Relative risk for a twofold increase in initial dioxin.

(d) MODEL 2: RANCH	HANDS – INITIAL DIOXIN – ADJUSTED	
	Analysis Results for Log ₂ (Initial Dioxin	n)
	Adjusted Relative Risk	
n	(95% C.I.) ^a	p-Value
457	1.00 (0.68,1.48)	0.981

^a Relative risk for a twofold increase in initial dioxin.

Table 14-18. Analysis of Arrhythmia (Continued)

(e) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY - UNADJUSTED

		Number (%)	Est. Relative Risk	
Dioxin Category	n	Yes	(95% C.I.) ^{ab}	p-Value
Comparison	1,195	65 (5.4)		
Background RH	376	18 (4.8)	0.90 (0.53,1.54)	0.703
Low RH	233	19 (8.2)	1.54 (0.90,2.61)	0.114
High RH	243	13 (5.3)	0.96 (0.52,1.77)	0.886
Low plus High RH	476	32 (6.7)	1.21 (0.77,1.88)	0.409

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: 1987 Dioxin ≤ 10 ppt.

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, 10 ppt < Initial Dioxin ≤ 94 ppt.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

(f) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY - ADJUSTED

		Adjusted Relative Risk	
Dioxin Category	n	" (95% C.I.) ^a	p-Value
Comparison	1,155		
Background RH	360	0.87 (0.49,1.57)	0.647
Low RH	221	1.17 (0.65,2.11)	0.596
High RH	236	1.10 (0.57,2.12)	0.774
Low plus High RH	457	1.13 (0.70,1.83)	0.604

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: $1987 \text{ Dioxin} \le 10 \text{ ppt.}$

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, $10 \text{ ppt} < \text{Initial Dioxin} \le 94 \text{ ppt}$.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

(g) MODEL 4: RANCH HANDS – 1987 DIOXIN – UNADJUSTED

1987 Dioxin Category Summary Statistics		Analysis Results for Log ₂ (1987 Dioxin + 1)		
1987 Dioxin	n	Number (%) Yes	Estimated Relative Risk (95% C.I.) ^a	p-Value
Low	284	14 (4.9)	0.99 (0.82,1.20)	0.932
Medium	281	20 (7.1)		
High	287	16 (5.6)		

^a Relative risk for a twofold increase in 1987 dioxin.

Note: Low = ≤ 7.9 ppt; Medium = > 7.9 - 19.6 ppt; High = > 19.6 ppt.

^b Adjusted for percent body fat at the time of the blood measurement of dioxin.

Table 14-18. Analysis of Arrhythmia (Continued)

(h) MODEL 4: RANCH HANDS – 1987 DIOXIN – ADJUSTED			
	Analysis Results for Log ₂ (1987 Dioxin + 1)		
	Adjusted Relative Risk		
n	(95% C.I.) ^a	p-Value	
817	1.12 (0.85,1.49)	0.422	

^a Relative risk for a twofold increase in 1987 dioxin.

14.2.2.2.13 Evidence of Prior Myocardial Infarction

The Model 1 unadjusted and adjusted analyses of prior myocardial infarction from the ECG showed no significant group differences over all participants or within each occupational stratum (Table 14-19(a,b): p>0.64 for each contrast).

Table 14-19. Analysis of Evidence of Prior Myocardial Infarction

Occupational Category	Group	n	Number (%) Yes	Est. Relative Risk (95% C.I.)	p-Value
All	Ranch Hand Comparison	859 1,232	34 (4.0) 53 (4.3)	0.92 (0.59,1.42)	0.698
Officer	Ranch Hand Comparison	334 484	15 (4.5) 23 (4.8)	0.94 (0.48,1.83)	0.862
Enlisted Flyer	Ranch Hand Comparison	149 186	7 (4.7) 9 (4.8)	0.97 (0.35,2.67)	0.952
Enlisted Groundcrew	Ranch Hand Comparison	376 562	12 (3.2) 21 (3.7)	0.85 (0.41,1.75)	0.657

(b) MODEL 1: RANCH HANDS VS. COMPARISONS – ADJUSTED						
Occupational Category	Adjusted Relative Risk (95% C.I.)	p-Value				
All	0.90 (0.56,1.43)	0.649				
Officer	0.88 (0.43,1.78)	0.718				
Enlisted Flyer	1.02 (0.35,2.96)	0.972				
Enlisted Groundcrew	0.86 (0.40,1.85)	0.709				

Table 14-19. Analysis of Evidence of Prior Myocardial Infarction (Continued)

(c) MODEL 2: RANCH HANDS – INITIAL DIOXIN – UNADJUSTED

(c) MODEL 2.		b Hilling Diomit		
Initial Dioxin Category Summary Statistics			Analysis Results for Log ₂ (Initial Dioxin) ^a
Initial Dioxin	n	Number (%) Yes	Estimated Relative Risk (95% C.I.) ^b	p-Value
Low	155	5 (3.2)	1.05 (0.75,1.46)	0.793
Medium	161	9 (5.6)		
High	160	7 (4.4)		

^a Adjusted for percent body fat at the time of the blood measurement of dioxin.

Note: Low = 27-63 ppt; Medium = >63-152 ppt; High = >152 ppt.

(d) MODEL 2: RANCH HANDS – INITIAL DIOXIN – ADJUSTED

(d) MODEL 2: KANCH HANDS	HATTIME DIOXIA ADSCRIED	
	Analysis Results for Log ₂ (Initial Diox	in)
	Adjusted Relative Risk	
n	(95% C.I.) ^a	p-Value
457	1.84 (1.13,2.99)	0.012

^a Relative risk for a twofold increase in initial dioxin.

Note: Results are not adjusted for race because of the sparse number of Ranch Hands with evidence of a prior myocardial infarction.

(e) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY – UNADJUSTED

		Number (%)	Est. Relative Risk	
Dioxin Category	n	Yes	(95% C.I.) ^{ab}	p-Value
Comparison	1,195	53 (4.4)		
Background RH	376	12 (3.2)	0.75 (0.39,1.42)	0.374
Low RH	233	11 (4.7)	1.06 (0.54,2.06)	0.867
High RH	243	10 (4.1)	0.88 (0.44,1.76)	0.722
Low plus High RH	476	21 (4.4)	0.96 (0.57,1.62)	0.891

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: 1987 Dioxin ≤ 10 ppt.

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, $10 \text{ ppt} < \text{Initial Dioxin} \le 94 \text{ ppt}$. High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

^b Relative risk for a twofold increase in initial dioxin.

b Adjusted for percent body fat at the time of the blood measurement of dioxin.

Table 14-19. Analysis of Evidence of Prior Myocardial Infarction (Continued)

(f) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY - ADJUSTED

		Adjusted Relative Risk	
Dioxin Category	n	(95% C.I.) ^a	p-Value
Comparison	1,155		
Background RH	360	0.69 (0.34,1.37)	0.285
Low RH	221	0.79 (0.39,1.61)	0.524
High RH	236	1.11 (0.52,2.36)	0.783
Low plus High RH	457	0.94 (0.54,1.65)	0.841

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: $1987 \text{ Dioxin} \le 10 \text{ ppt.}$

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, $10 \text{ ppt} < \text{Initial Dioxin} \le 94 \text{ ppt}$. High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

(g) MODEL 4.	RANCH HANDS -	1987 DIOXIN -	- IINADIIISTED
(E) MODEL T.	MANUH HANDS =	170/ DIOAII1 =	

(g) 1.10222 IV		.20 20.210111		
1987 Dioxin Category Summary Statistics			Analysis Results for Log ₂	(1987 Dioxin + 1)
1987 Dioxin	n	Number (%) Yes	Estimated Relative Risk (95% C.I.) ^a	p-Value
Low	284	7 (2.5)	1.09 (0.87,1.38)	0.447
Medium	281	12 (4.3)		
High	287	14 (4.9)		

^a Relative risk for a twofold increase in 1987 dioxin.

Note: Low = ≤ 7.9 ppt; Medium = > 7.9 - 19.6 ppt; High = > 19.6 ppt.

(h) MODEL 4: RANCH HANDS – 1987 DIOXIN – ADJUSTED						
	Analysis Results for Log ₂ (1987 Dioxin + 1)					
	Adjusted Relative Risk					
n	(95% C.I.) ^a	p-Value				
817	1.33 (0.95,1.87)	0.089				

^a Relative risk for a twofold increase in 1987 dioxin.

The unadjusted Model 2 analysis revealed no significant association between initial dioxin and prior myocardial infarction (Table 14-19(c): p=0.793). After adjusting for covariates, the results became significant (Table 14-19(d): Adj. RR=1.84, p=0.012). The percentages of participants with evidence of prior myocardial infarction in the low, medium, and high initial dioxin categories were 3.2, 5.6, and 4.4, respectively.

The unadjusted and adjusted Model 3 analyses of prior myocardial infarction did not show any of the Ranch Hand categories to be significantly different from the Comparisons (Table 14-19(e,f): p>0.28 for each contrast).

The unadjusted Model 4 analysis revealed no significant association between 1987 dioxin and evidence of prior myocardial infarction (Table 14-19(g): p=0.447). After adjusting for covariates, the results became marginally significant (Table 14-19(h): Adj. RR=1.33, p=0.089). The percentages of participants with evidence of prior myocardial infarction in the low, medium, and high 1987 dioxin categories were 2.5, 4.3, and 4.9, respectively.

14.2.2.2.14 ECG: Other Diagnoses

The Model 1 unadjusted and adjusted analyses of other ECG diagnoses showed no significant group differences over all participants or within each occupational stratum (Table 14-20(a,b): p>0.15 for each contrast).

Table 14-20. Analysis of ECG: Other Diagnoses

Occupational	G		Number (%)	Est. Relative Risk	¥7. 1
Category	Group	n	Yes	(95% C.I.)	p-Value
All	Ranch Hand	859	3 (0.3)	4.31 (0.45,41.55)	0.168
	Comparison	1,232	1 (0.1)		
Officer	Ranch Hand	334	1 (0.3)		0.852^{a}
	Comparison	484	0 (0.0)		
Enlisted Flyer	Ranch Hand	149	0 (0.0)		
•	Comparison	186	0 (0.0)		
Enlisted	Ranch Hand	376	2 (0.5)	3.00 (0.27,33.20)	0.370
Groundcrew	Comparison	562	1 (0.2)		

^a P-value determined using a chi-square test with continuity correction because of the sparse number of participants with other abnormal ECG diagnoses.

^{--:} Results not presented because of the sparse number of participants with other abnormal ECG diagnoses.

(b) MODEL 1: RANCH HANDS VS. COMPARISONS – ADJUSTED						
Occupational Category	Adjusted Relative Risk (95% C.I.)	p-Value				
All	4.67 (0.47,46.79)	0.153				
Officer Enlisted Flyer	 	 				
Enlisted Groundcrew	3.29 (0.28,38.94)	0.346				

^{--:} Results not presented because of the sparse number of participants with other abnormal ECG diagnoses.

Note: Results are not adjusted for family history of heart disease before age 45 and diabetic class because of the sparse number of participants with other abnormal ECG diagnoses. Results for all occupations combined also are not adjusted for occupation because of the sparse number of participants with other abnormal ECG diagnoses.

Table 14-20. Analysis of ECG: Other Diagnoses (Continued)

(c) MODEL 2: RANCH HANDS – INITIAL DIOXIN – UNADJUSTED

Initial Dioxin Category Summary Statistics			Analysis Results for Log ₂ (Initial Dioxin) ^a
Initial Dioxin	n	Number (%) Yes	Estimated Relative Risk (95% C.I.) ^b	p-Value
Low	155	0 (0.0)	1.53 (0.62,3.79)	0.381
Medium	161	0 (0.0)		
High	160	2 (1.3)		

^a Adjusted for percent body fat at the time of the blood measurement of dioxin.

Note: Low = 27-63 ppt; Medium = >63-152 ppt; High = >152 ppt.

(d) MODEL 2: RANCH HANDS – INITIAL DIOXIN – ADJUSTED

(a) MODEL 2: RANCH HANDS - INITIAL DIOXIN - ADJUSTED				
Analysis Results for Log ₂ (Initial Dioxin)				
Adjusted Relative Risk				
n	(95% C.I.)	p-Value		

--: Results not presented because of the sparse number of Ranch Hands with other abnormal ECG diagnoses.

(e) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY – UNADJUSTED

		Number (%)	Est. Relative Risk	
Dioxin Category	n	Yes	(95% C.I.) ^{ab}	p-Value
Comparison	1,195	1 (0.1)		
Background RH	376	1 (0.3)	2.59 (0.16,41.85)	0.503
Low RH	233	0(0.0)		0.999^{c}
High RH	243	2 (0.8)	12.49 (1.10,142.56)	0.042
Low plus High RH	476	2 (0.4)		0.409^{c}

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: 1987 Dioxin ≤ 10 ppt.

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, $10 \text{ ppt} < \text{Initial Dioxin} \le 94 \text{ ppt}$.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

^b Relative risk for a twofold increase in initial dioxin.

^b Adjusted for percent body fat at the time of the blood measurement of dioxin.

^c P-value determined using a chi-square test with continuity correction because of the sparse number of participants with other abnormal ECG diagnoses.

^{--:} Results not presented because of the sparse number of participants with other abnormal ECG diagnoses.

Table 14-20. Analysis of ECG: Other Diagnoses (Continued)

(f) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY - ADJUSTED

		Adjusted Relative Risk	
Dioxin Category	n	(95% C.I.) ^a	p-Value
Comparison	1,186		
Background RH	368	2.89 (0.16,52.97)	0.474
Low RH	227		
High RH	239	12.41 (1.00,154.15)	0.050
Low plus High RH	466		

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: $1987 \text{ Dioxin} \le 10 \text{ ppt.}$

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, 10 ppt < Initial Dioxin ≤ 94 ppt.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

Results are not adjusted for occupation, family history of heart disease before age 45, and diabetic class because of the sparse number of participants with other abnormal ECG diagnoses.

(g) MODEL 4: RANCH HANDS – 1987 DIOXIN – UNADJUSTED

1987 Dioxin Category Summary Statistics			Analysis Results for Log ₂	(1987 Dioxin + 1)
1987 Dioxin	n	Number (%) Yes	Estimated Relative Risk (95% C.I.) ^a p-Value	
Low	284	1 (0.4)	1.27 (0.63,2.59)	0.512
Medium	281	0 (0.0)		
High	287	2 (0.7)		

^a Relative risk for a twofold increase in 1987 dioxin.

Note: Low = \le 7.9 ppt; Medium = >7.9–19.6 ppt; High = >19.6 ppt.

Analysis Results for Log ₂ (1987 Dioxin + 1)					
	Adjusted Relative Risk				
n	(95% C.I.) ^a	p-Value			
834	1.47 (0.58,3.73)	0.413			

^a Relative risk for a twofold increase in 1987 dioxin.

Note: Results are not adjusted for occupation, current cigarette smoking, family history of heart disease before age 45, and diabetic class because of the sparse number of Ranch Hands with other abnormal ECG diagnoses.

^{--:} Results not presented because of the sparse number of participants with other abnormal ECG diagnoses.

The unadjusted Model 2 analysis revealed no significant results (Table 14-20(c): p=0.381). Because of the sparse number of Ranch Hands with other ECG diagnoses, the adjusted Model 2 analysis was not performed.

The unadjusted and adjusted Model 3 analyses each revealed a significant difference between Ranch Hands in the high dioxin category and Comparisons (Table 14-20(e,f): Est. RR=12.49, p=0.042; Adj. RR=12.41, p=0.050, respectively). The percentage of Ranch Hands in the high dioxin category was 0.8 versus 0.1 percent for the Comparisons.

Both the unadjusted and adjusted Model 4 analyses did not reveal a significant association between 1987 dioxin and other ECG diagnoses (Table 14-20(g,h): p<0.41 for each analysis).

14.2.2.3 Physical Examination Variables – Peripheral Vascular Function

14.2.2.3.1 Funduscopic Examination

The unadjusted and adjusted Model 1 analyses of funduscopic examination did not reveal a group difference between Ranch Hands and Comparisons when all occupations were combined (Table 14-21(a,b): p>0.56 for each contrast). Stratifying by occupation revealed a significant group difference within the enlisted groundcrew stratum in both the unadjusted and adjusted analyses (Table 14-21(a,b): Est. RR=0.62, p=0.033; Adj. RR=0.62, p=0.047, respectively). Ranch Hand enlisted groundcrew had fewer abnormal funduscopic examination results (8.8%) than did Comparison enlisted groundcrew (13.3%).

Table 14-21. Analysis of Funduscopic Examination

(a) MODEL 1: RANCH HANDS VS. COMPARISONS – UNADJUSTED					
Occupational Category	Group	n	Number (%) Abnormal	Est. Relative Risk (95% C.I.)	p-Value
All	Ranch Hand Comparison	858 1,231	105 (12.2) 156 (12.7)	0.96 (0.74,1.25)	0.767
Officer	Ranch Hand Comparison	333 484	42 (12.6) 49 (10.1)	1.28 (0.83,1.99)	0.267
Enlisted Flyer	Ranch Hand Comparison	149 185	30 (20.1) 32 (17.3)	1.21 (0.69,2.09)	0.508
Enlisted Groundcrew	Ranch Hand Comparison	376 562	33 (8.8) 75 (13.3)	0.62 (0.41,0.96)	0.033

(b) MODEL 1: RANCH HANDS VS. COMPARISONS – ADJUSTED					
Occupational Category	Adjusted Relative Risk (95% C.I.)	p-Value			
All	0.92 (0.69,1.22)	0.562			
Officer	1.27 (0.79,2.02)	0.321			
Enlisted Flyer	1.06 (0.59,1.91)	0.852			
Enlisted Groundcrew	0.62 (0.39,0.99)	0.047			

Table 14-21. Analysis of Funduscopic Examination (Continued)

(c) MODEL 2: RANCH HANDS – INITIAL DIOXIN – UNADJUSTED					
Initial I	Dioxin Category S	Summary Statistics	Analysis Results for Log ₂ (Initial Dioxin) ^a	
Initial Dioxin	n	Number (%) Abnormal	Estimated Relative Risk (95% C.I.) ^b	p-Value	
Low	155	20 (12.9)	0.93 (0.76,1.15)	0.520	
Medium	161	24 (14.9)			
High	160	18 (11 3)			

^a Adjusted for percent body fat at the time of the blood measurement of dioxin.

Note: Low = 27-63 ppt; Medium = >63-152 ppt; High = >152 ppt.

(d) MODEL 2: RANCH HANDS – INITIAL DIOXIN – ADJUSTED					
Analysis Results for Log ₂ (Initial Dioxin)					
	Adjusted Relative Risk				
n	(95% C.I.) ^a	p-Value			
457	1.14 (0.87,1.50)	0.342			

^a Relative risk for a twofold increase in initial dioxin.

(e) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY – UNADJUSTED

		Number (%)	Est. Relative Risk	
Dioxin Category	n	Abnormal	(95% C.I.) ^{ab}	p-Value
Comparison	1,194	149 (12.5)		
Background RH	375	43 (11.5)	0.99 (0.69,1.43)	0.963
Low RH	233	30 (12.9)	1.02 (0.67,1.56)	0.921
High RH	243	32 (13.2)	0.98 (0.65,1.49)	0.933
Low plus High RH	476	62 (13.0)	1.00 (0.73,1.38)	0.993

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: 1987 Dioxin ≤ 10 ppt.

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, $10 \text{ ppt} < \text{Initial Dioxin} \le 94 \text{ ppt}$.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

^b Relative risk for a twofold increase in initial dioxin.

^b Adjusted for percent body fat at the time of the blood measurement of dioxin.

Table 14-21. Analysis of Funduscopic Examination (Continued)

(f) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY - ADJUSTED

		Adjusted Relative Risk	
Dioxin Category	n	(95% C.I.) ^a	p-Value
Comparison	1,154		
Background RH	359	1.04 (0.70,1.55)	0.842
Low RH	221	0.82 (0.52,1.30)	0.402
High RH	236	0.95 (0.60,1.51)	0.836
Low plus High RH	457	0.89 (0.63,1.26)	0.500

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: 1987 Dioxin ≤ 10 ppt.

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, $10 \text{ ppt} < \text{Initial Dioxin} \le 94 \text{ ppt}$. High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

(g) MODEL 4: RANCH HANDS – 1987 DIOXIN – UNADJUSTED

(3)						
1987 Dioxin Category Summary Statistics		Analysis Results for Log ₂ (1987 Dioxin + 1)				
1987 Dioxin	n	Number (%) Abnormal	Estimated Relative Risk (95% C.I.) ^a	p-Value		
Low	283	30 (10.6)	1.00 (0.87,1.15)	0.951		
Medium	281	36 (12.8)				
High	287	39 (13.6)				

^a Relative risk for a twofold increase in 1987 dioxin.

Note: Low = \le 7.9 ppt; Medium = >7.9–19.6 ppt; High = >19.6 ppt.

(h) MODEL 4: RANCH HANDS – 1987 DIOXIN – ADJUSTED					
Analysis Results for Log ₂ (1987 Dioxin + 1)					
	Adjusted Relative Risk				
n	(95% C.I.) ^a	p-Value			
816	1.03 (0.85,1.24)	0.767			

^a Relative risk for a twofold increase in 1987 dioxin.

The unadjusted and adjusted analyses in Models 2 through 4 were nonsignificant (Table 14-21(c-h): p>0.34 for each analysis).

14.2.2.3.2 Carotid Bruits

All Model 1 through 4 unadjusted and adjusted analyses were nonsignificant (Table 14-22(a–h): p>0.21 for each analysis).

Table 14-22. Analysis of Carotid Bruits

(a) MODEL	1.	RANCH H	ANDS VS.	COMPARISONS	- UNADJUSTED
			7171717 V 17.		

Occupational Category	Group	n	Number (%) Abnormal	Est. Relative Risk (95% C.I.)	p-Value
All	Ranch Hand Comparison	859 1,232	23 (2.7) 33 (2.7)	1.00 (0.58,1.71)	0.999
Officer	Ranch Hand Comparison	334 484	6 (1.8) 12 (2.5)	0.72 (0.27,1.94)	0.515
Enlisted Flyer	Ranch Hand Comparison	149 186	8 (5.4) 5 (2.7)	2.05 (0.66,6.41)	0.215
Enlisted Groundcrew	Ranch Hand Comparison	376 562	9 (2.4) 16 (2.8)	0.84 (0.37,1.91)	0.673

(b) MODEL 1: RANCH HANDS VS. COMPARISONS – ADJUSTED

	Adjusted Relative Risk	
Occupational Category	(95% C.I.)	p-Value
All	0.94 (0.53,1.65)	0.823
Officer	0.72 (0.26,1.99)	0.524
Enlisted Flyer	1.94 (0.58,6.46)	0.283
Enlisted Groundcrew	0.78 (0.33,1.86)	0.578

(c) MODEL 2: RANCH HANDS – INITIAL DIOXIN – UNADJUSTED

Initial Dioxin Category Summary Statistics			Analysis Results for Log ₂ (Initial Dioxin) ^a
Initial Dioxin	n	Number (%) Abnormal	Estimated Relative Risk (95% C.I.) ^b	p-Value
Low	155	3 (1.9)	1.06 (0.70,1.59)	0.797
Medium	161	5 (3.1)		
High	160	5 (3.1)		

^a Adjusted for percent body fat at the time of the blood measurement of dioxin. ^b Relative risk for a twofold increase in initial dioxin.

Note: Low = 27-63 ppt; Medium = >63-152 ppt; High = >152 ppt.

(d) MODEL 2: RANCH HANDS – INITIAL DIOXIN – ADJUSTED					
	Analysis Results for Log ₂ (Initial Dioxin	n)			
	Adjusted Relative Risk				
n	(95% C.I.) ^a	p-Value			
457	1.15 (0.62,2.11)	0.658			

^a Relative risk for a twofold increase in initial dioxin.

Table 14-22. Analysis of Carotid Bruits (Continued)

(e) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY - UNADJUSTED

		Number (%)	Est. Relative Risk	
Dioxin Category	n	Abnormal	(95% C.I.) ^{ab}	p-Value
Comparison	1,195	31 (2.6)		
Background RH	376	9 (2.4)	0.93 (0.44,1.98)	0.853
Low RH	233	5 (2.1)	0.82 (0.32,2.14)	0.687
High RH	243	8 (3.3)	1.27 (0.57,2.80)	0.561
Low plus High RH	476	13 (2.7)	1.02 (0.53,2.00)	0.943

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: 1987 Dioxin ≤ 10 ppt.

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, 10 ppt < Initial Dioxin ≤ 94 ppt.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

(f) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY - ADJUSTED

		Adjusted Relative Risk	
Dioxin Category	n	(95% C.I.) ^a	p-Value
Comparison	1,155		
Background RH	360	1.06 (0.47,2.38)	0.893
Low RH	221	0.69 (0.25,1.86)	0.460
High RH	236	1.01 (0.41,2.45)	0.991
Low plus High RH	457	0.84 (0.41,1.71)	0.625

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: $1987 \text{ Dioxin} \le 10 \text{ ppt.}$

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, $10 \text{ ppt} < \text{Initial Dioxin} \le 94 \text{ ppt}$.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

(g) MODEL 4: RANCH HANDS – 1987 DIOXIN – UNADJUSTED

1987 Dioxin Category Summary Statistics			Analysis Results for Log ₂	(1987 Dioxin + 1)
1987 Dioxin	n	Number (%) Abnormal	Estimated Relative Risk (95% C.I.) ^a	p-Value
Low	284	7 (2.5)	1.02 (0.77,1.36)	0.897
Medium	281	7 (2.5)		
High	287	8 (2.8)		

^a Relative risk for a twofold increase in 1987 dioxin.

Note: Low = ≤ 7.9 ppt; Medium = > 7.9 - 19.6 ppt; High = > 19.6 ppt.

^b Adjusted for percent body fat at the time of the blood measurement of dioxin.

Table 14-22. Analysis of Carotid Bruits (Continued)

(h) MODEL 4: RANCH HANDS – 1987 DIOXIN – ADJUSTED						
	Analysis Results for Log ₂ (1987 Dioxin + 1)					
	Adjusted Relative Risk					
n	(95% C.I.) ^a	p-Value				
817	0.94 (0.65,1.36)	0.755				

^a Relative risk for a twofold increase in 1987 dioxin.

14.2.2.3.3 Radial Pulses

The unadjusted and adjusted Model 1 analyses of radial pulses were nonsignificant (Table 14-23(a,b): p>0.11 for each contrast).

Table 14-23. Analysis of Radial Pulses

Occupational Category	Group	n	Number (%) Abnormal	Est. Relative Risk (95% C.I.)	p-Value
All	Ranch Hand Comparison	859 1,232	7 (0.8) 4 (0.3)	2.52 (0.74,8.64)	0.131
Officer	Ranch Hand Comparison	334 484	2 (0.6) 2 (0.4)	1.45 (0.20,10.36)	0.710
Enlisted Flyer	Ranch Hand Comparison	149 186	0 (0.0) 0 (0.0)		
Enlisted Groundcrew	Ranch Hand Comparison	376 562	5 (1.3) 2 (0.4)	3.77 (0.73,19.55)	0.114

^{--:} Results not presented because of the sparse number of participants with an abnormal radial pulse.

(b) MODEL 1: RANCH HANDS VS. COMPARISONS – ADJUSTED					
Occupational Category	Adjusted Relative Risk (95% C.I.)	p-Value			
All	2.85 (0.67,12.16)	0.143			
Officer Enlisted Flyer Enlisted Groundcrew	1.24 (0.16,9.95) 5.69 (0.54,60.05)	0.837 0.148			

^{--:} Results not presented because of the sparse number of participants with an abnormal radial pulse.

Note: Results for all occupations combined are not adjusted for occupation because of the sparse number of participants with an abnormal radial pulse.

Table 14-23. Analysis of Radial Pulses (Continued)

(c) MODEL 2: RANCH HANDS – INITIAL DIOXIN – UNADJUSTED

Initial Dioxin Category Summary Statistics			Analysis Results for Log ₂ (Initial Dioxin) ^a		
Initial Dioxin	n	Number (%) Abnormal	Estimated Relative Risk (95% C.I.) ^b	p-Value	
Low	155	2 (1.3)	0.58 (0.17,1.99)	0.334	
Medium	161	0 (0.0)			
High	160	1 (0.6)			

^a Adjusted for percent body fat at the time of the blood measurement of dioxin.

Note: Low = 27-63 ppt; Medium = >63-152 ppt; High = >152 ppt.

(d) MODEL 2: RANCH HANDS – INITIAL DIOXIN – ADJUSTED

(u) MODEL 2: KANCH I	(u) MODEL 2: KANCH HANDS - INITIAL DIOXIN - ADJUSTED			
	Analysis Results for Log ₂ (Initial Dioxi	n)		
	Adjusted Relative Risk			
n	(95% C.I.)	p-Value		

^{--:} Results not presented because of the sparse number of Ranch Hands with an abnormal radial pulse.

(e) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY – UNADJUSTED

		Number (%)	Est. Relative Risk	
Dioxin Category	n	Abnormal	(95% C.I.) ^{ab}	p-Value
Comparison	1,195	4 (0.3)		
Background RH	376	4 (1.1)	2.78 (0.69,11.27)	0.153
Low RH	233	2 (0.9)	2.64 (0.48,14.54)	0.264
High RH	243	1 (0.4)	1.41 (0.16,12.80)	0.759
Low plus High RH	476	3 (0.6)	1.92 (0.40,9.18)	0.414

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: 1987 Dioxin ≤ 10 ppt.

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, $10 \text{ ppt} < \text{Initial Dioxin} \le 94 \text{ ppt}$.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

^b Relative risk for a twofold increase in initial dioxin.

^b Adjusted for percent body fat at the time of the blood measurement of dioxin.

Table 14-23. Analysis of Radial Pulses (Continued)

(f) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY - ADJUSTED

		A diseased Delection Diele	
Dioxin Category	n	Adjusted Relative Risk (95% C.I.) ^a	p-Value
Comparison	1,155		
Background RH	360	3.27 (0.64,16.71)	0.155
Low RH	221	3.82 (0.53,27.51)	0.183
High RH	236	1.26 (0.11,14.89)	0.856
Low plus High RH	457	2.15 (0.36,13.04)	0.404

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: $1987 \text{ Dioxin} \le 10 \text{ ppt.}$

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, $10 \text{ ppt} < \text{Initial Dioxin} \le 94 \text{ ppt}$.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

Results are not adjusted for occupation because of the sparse number of participants with an abnormal radial pulse.

		~	~	400	
(g) MODEL	4:	RANCH	HANDS -	· 1987 DIOXIN =	UNADJUSTED

1987 Dioxin Category Summary Statistics			Analysis Results for Log ₂	(1987 Dioxin + 1)
1987 Dioxin	n	Number (%) Abnormal	Estimated Relative Risk (95% C.I.) ^a	p-Value
Low	284	2 (0.7)	0.75 (0.43,1.32)	0.305
Medium	281	4 (1.4)		
High	287	1 (0.3)		

^a Relative risk for a twofold increase in 1987 dioxin.

Note: Low = ≤ 7.9 ppt; Medium = > 7.9 - 19.6 ppt; High = > 19.6 ppt.

(h)	MODEL	4:	RANCH	HANDS -	1987	DIOXIN -	ADJUSTED

Analysis Results for Log ₂ (1987 Dioxin + 1)				
n	Adjusted Relative Risk (95% C.I.) ^a	p-Value		
817	0.61 (0.30,1.21)	0.140		

^a Relative risk for a twofold increase in 1987 dioxin.

Note: Results are not adjusted for occupation because of the sparse number of Ranch Hands with an abnormal radial pulse.

The unadjusted Model 2 analysis showed no significant association between radial pulses and initial dioxin (Table 14-23(c): p=0.334). Because of the sparse number of Ranch Hands with abnormal radial pulses, the adjusted Model 2 analysis was not performed.

All Model 3 and 4 analyses of radial pulses were nonsignificant (Table 14-23(e−h): p≥0.14 for each analysis).

14.2.2.3.4 Femoral Pulses

The unadjusted Model 1 analysis of femoral pulses revealed a marginally significant overall group difference between Ranch Hands and Comparisons (Table 14-24(a): Est. RR=1.83, p=0.080). Stratifying by occupation did not reveal any significant difference between Ranch Hands and Comparisons within each occupational stratum (Table 14-24(a): p>0.12 for each contrast). The percentage of participants with abnormal femoral pulses was greater for the Ranch Hands (2.2%) than for Comparisons (1.2%). The adjusted analysis did not show a significant difference between Ranch Hands and Comparisons over all occupations or within each occupational stratum (Table 14-24(b): p>0.17 for each contrast).

Table 14-24. Analysis of Femoral Pulses

(a) MODEL 1:	(a) MODEL 1: RANCH HANDS VS. COMPARISONS – UNADJUSTED				
Occupational Category	Group	n	Number (%) Abnormal	Est. Relative Risk (95% C.I.)	p-Value
All	Ranch Hand Comparison	859 1,231	19 (2.2) 15 (1.2)	1.83 (0.93,3.63)	0.080
Officer	Ranch Hand Comparison	334 484	7 (2.1) 8 (1.7)	1.27 (0.46,3.55)	0.643
Enlisted Flyer	Ranch Hand Comparison	149 185	5 (3.4) 3 (1.6)	2.11 (0.50,8.96)	0.313
Enlisted Groundcrew	Ranch Hand Comparison	376 562	7 (1.9) 4 (0.7)	2.65 (0.77,9.10)	0.123

(b) MODEL 1: RANCH HANDS VS. COMPARISONS – ADJUSTED				
Occupational Category	Adjusted Relative Risk (95% C.I.)	p-Value		
All	1.66 (0.79,3.49)	0.178		
Officer	1.51 (0.52,4.38)	0.448		
Enlisted Flyer	1.48 (0.27,8.02)	0.652		
Enlisted Groundcrew	2.08 (0.55,7.87)	0.282		

(c) MODEL 2:	(c) MODEL 2: RANCH HANDS – INITIAL DIOXIN – UNADJUSTED				
Initial I	Initial Dioxin Category Summary Statistics Analysis Results for Log ₂ (Initial Dioxin) ^a				
Initial Dioxin	n	Number (%) Abnormal	Estimated Relative Risk (95% C.I.) ^b	p-Value	
Low	155	3 (1.9)	0.97 (0.61,1.53)	0.890	
Medium	161	5 (3.1)			
High	160	4 (2.5)			

^a Adjusted for percent body fat at the time of the blood measurement of dioxin.

Note: Low = 27-63 ppt; Medium = >63-152 ppt; High = >152 ppt.

^b Relative risk for a twofold increase in initial dioxin.

Table 14-24. Analysis of Femoral Pulses (Continued)

(d) MODEL 2: RANCH I	(d) MODEL 2: RANCH HANDS – INITIAL DIOXIN – ADJUSTED			
	Analysis Results for Log ₂ (Initial Dioxi	(n)		
	Adjusted Relative Risk			
n	(95% C.I.) ^a	p-Value		
457	1 17 (0 61 2 24)	0.641		

^a Relative risk for a twofold increase in initial dioxin.

Note: Results are not adjusted for race because of the sparse number of participants with an abnormal femoral pulse.

(e) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY - UNADJUSTED

		Number (%)	Est. Relative Risk	
Dioxin Category	n	Abnormal	(95% C.I.) ^{ab}	p-Value
Comparison	1,194	15 (1.3)		
Background RH	376	7 (1.9)	1.39 (0.56,3.45)	0.481
Low RH	233	6 (2.6)	2.10 (0.81,5.48)	0.128
High RH	243	6 (2.5)	2.13 (0.81,5.56)	0.125
Low plus High RH	476	12 (2.5)	2.11 (0.98,4.56)	0.056

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: 1987 Dioxin ≤ 10 ppt.

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, 10 ppt < Initial Dioxin ≤ 94 ppt.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

(f) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY - ADJUSTED

		Adjusted Relative Risk	
Dioxin Category	n	(95% C.I.) ^a	p-Value
Comparison	1,154		
Background RH	360	1.22 (0.44,3.36)	0.702
Low RH	221	1.71 (0.58,4.98)	0.329
High RH	236	2.45 (0.76,7.90)	0.134
Low plus High RH	457	2.06 (0.85,4.96)	0.108

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: $1987 \text{ Dioxin} \le 10 \text{ ppt.}$

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, $10 \text{ ppt} < \text{Initial Dioxin} \le 94 \text{ ppt}$.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

^b Adjusted for percent body fat at the time of the blood measurement of dioxin.

Table 14-24. Analysis of Femoral Pulses (Continued)

(g) MODEL 4: RANCH HANDS – 1987 DIOXIN – UNADJUSTED						
1987 Dioxin Category Summary Statistics			Analysis Results for Log ₂	(1987 Dioxin + 1)		
1987 Dioxin	n	Number (%) Abnormal	Estimated Relative Risk (95% C.I.) ^a	p-Value		
Low	284	5 (1.8)	1.01 (0.75,1.38)	0.927		
Medium	281	5 (1.8)				
High	287	9 (3.1)				

^a Relative risk for a twofold increase in 1987 dioxin.

Note: Low = \le 7.9 ppt; Medium = >7.9–19.6 ppt; High = >19.6 ppt.

(h) MODEL 4: RANCH HANDS – 1987 DIOXIN – ADJUSTED					
	Analysis Results for Log ₂ (1987 Dioxin + 1)				
	Adjusted Relative Risk				
n	(95% C.I.) ^a	p-Value			
817	1.29 (0.83,2.03)	0.255			

^a Relative risk for a twofold increase in 1987 dioxin.

The Model 2 analyses did not reveal a significant association between femoral pulses and initial dioxin in either the unadjusted or adjusted analyses (Table 14-24(c,d): p>0.64 for each analysis).

The unadjusted Model 3 analysis showed a marginally significant difference between Ranch Hands in the low plus high dioxin category and Comparisons (Table 14-24(e): Est. RR=2.11, p=0.056). The percentage of abnormal femoral pulses for Ranch Hands in the low plus high dioxin category was 2.5 versus 1.3 percent for Comparisons. The adjusted analysis did not find any contrasts to be significant (Table 14-24(f): p>0.10 for each contrast).

The unadjusted and adjusted Model 4 analyses did not show a significant association between 1987 dioxin and femoral pulses (Table 14-24(g,h): p>0.25 for each analysis).

14.2.2.3.5 Popliteal Pulses

All unadjusted and adjusted Model 1 through 4 analyses were not significant (Table 14-25(a−h): p≥0.41 for each analysis).

Table 14-25. Analysis of Popliteal Pulses

(a) MODEL 1: RANCH HANDS VS. COMPARISONS – UNADJUSTED					
Occupational Category	Group	n	Number (%) Abnormal	Est. Relative Risk (95% C.I.)	p-Value
All	Ranch Hand Comparison	859 1,230	23 (2.7) 28 (2.3)	1.18 (0.68,2.06)	0.561
Officer	Ranch Hand Comparison	334 483	7 (2.1) 12 (2.5)	0.84 (0.33,2.16)	0.717
Enlisted Flyer	Ranch Hand Comparison	149 185	5 (3.4) 4 (2.2)	1.57 (0.41,5.96)	0.506
Enlisted Groundcrew	Ranch Hand Comparison	376 562	11 (2.9) 12 (2.1)	1.38 (0.60,3.16)	0.445

Table 14-25. Analysis of Popliteal Pulses (Continued)

(b) MODEL 1: RANCH HANDS VS. COMPARISONS – ADJUSTED

(b) MODEL 1. REPORTED VO. COMPTRISONS INDUCTIES						
Occupational Category	Adjusted Relative Risk (95% C.I.)	p-Value				
All	1.04 (0.56,1.90)	0.911				
Officer	0.95 (0.35,2.52)	0.911				
Enlisted Flyer	0.99 (0.21,4.82)	0.995				
Enlisted Groundcrew	1.13 (0.46,2.79)	0.784				

(c) MODEL 2: RANCH HANDS - INITIAL DIOXIN - UNADJUSTED

Initial Dioxin Category Summary Statistics			Analysis Results for Log ₂ (Initial Dioxin) ^a
Initial Dioxin	n	Number (%) Abnormal	Estimated Relative Risk (95% C.I.) ^b	p-Value
Low	155	4 (2.6)	0.89 (0.57,1.38)	0.601
Medium	161	6 (3.7)		
High	160	4 (2.5)		

^a Adjusted for percent body fat at the time of the blood measurement of dioxin.

Note: Low = 27-63 ppt; Medium = >63-152 ppt; High = >152 ppt.

(d) MODEL 2: RANCH HANDS – INITIAL DIOXIN – ADJUSTED

()						
Analysis Results for Log ₂ (Initial Dioxin)						
Adjusted Relative Risk						
n	(95% C.I.) ^a	p-Value				
457	0.97 (0.53,1.78)	0.924				

^a Relative risk for a twofold increase in initial dioxin.

Note: Results are not adjusted for race because of the sparse number of Ranch Hands with an abnormal popliteal pulse.

(e) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY – UNADJUSTED

Dioxin Category	n	Number (%) Abnormal	Est. Relative Risk (95% C.I.) ^{ab}	p-Value
Comparison	1,193	28 (2.3)		<u>. </u>
Background RH	376	9 (2.4)	0.94 (0.44,2.03)	0.879
Low RH	233	7 (3.0)	1.31 (0.56,3.03)	0.535
High RH	243	7 (2.9)	1.33 (0.57,3.08)	0.512
Low plus High RH	476	14 (2.9)	1.32 (0.69,2.53)	0.410

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: $1987 \text{ Dioxin} \le 10 \text{ ppt.}$

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, $10 \text{ ppt} < \text{Initial Dioxin} \le 94 \text{ ppt}$.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

^b Relative risk for a twofold increase in initial dioxin.

^b Adjusted for percent body fat at the time of the blood measurement of dioxin.

Table 14-25. Analysis of Popliteal Pulses (Continued)

(f) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY - ADJUSTED

		Adjusted Relative Risk	
Dioxin Category	n	(95% C.I.) ^a	p-Value
Comparison	1,153		
Background RH	360	0.88 (0.37,2.05)	0.760
Low RH	221	1.15 (0.45,2.92)	0.776
High RH	236	1.08 (0.40,2.86)	0.884
Low plus High RH	457	1.11 (0.53,2.30)	0.781

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: 1987 Dioxin ≤ 10 ppt.

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, 10 ppt < Initial Dioxin ≤ 94 ppt. High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

(g) MODEL 4: RANCH HANDS – 1987 DIOXIN – UNADJUSTED

(8)							
1987 Dioxin Category Summary Statistics		Analysis Results for Log ₂	(1987 Dioxin + 1)				
1987 Number (%) Dioxin n Abnormal		Estimated Relative Risk (95% C.I.) ^a	p-Value				
Low	284	6 (2.1)	0.98 (0.74,1.30)	0.891			
Medium	281	7 (2.5)					
High	287	10 (3.5)					

^a Relative risk for a twofold increase in 1987 dioxin.

Note: Low = \le 7.9 ppt; Medium = >7.9–19.6 ppt; High = >19.6 ppt.

(h) MODEL 4: RANCH HANDS – 1987 DIOXIN – ADJUSTED					
Analysis Results for Log ₂ (1987 Dioxin + 1)					
	Adjusted Relative Risk				
n	(95% C.I.) ^a	p-Value			
817	1.02 (0.72,1.46)	0.908			

^a Relative risk for a twofold increase in 1987 dioxin.

14.2.2.3.6 Dorsalis Pedis Pulses

All unadjusted and adjusted analyses of dorsalis pedis pulses were nonsignificant (Table 14-26(a–h): p>0.11 for each analysis).

Table 14-26. Analysis of Dorsalis Pedis Pulses

(a) MODEL 1:	RANCH HANDS VS.	COMPARISONS -	UNADJUSTED

Occupational Category	Group	n	Number (%) Abnormal	Est. Relative Risk (95% C.I.)	p-Value
All	Ranch Hand Comparison	859 1,230	69 (8.0) 95 (7.7)	1.04 (0.76,1.44)	0.796
Officer	Ranch Hand Comparison	334 483	27 (8.1) 32 (6.6)	1.24 (0.73,2.11)	0.429
Enlisted Flyer	Ranch Hand Comparison	149 185	18 (12.1) 17 (9.2)	1.36 (0.67,2.74)	0.392
Enlisted Groundcrew	Ranch Hand Comparison	376 562	24 (6.4) 46 (8.2)	0.76 (0.46,1.28)	0.305

(b) MODEL 1: RANCH HANDS VS. COMPARISONS – ADJUSTED

	Adjusted Relative Risk	
Occupational Category	(95% C.I.)	p-Value
All	0.97 (0.69,1.37)	0.857
Officer	1.27 (0.73,2.22)	0.398
Enlisted Flyer	1.33 (0.62,2.86)	0.463
Enlisted Groundcrew	0.64 (0.37,1.12)	0.117

(c) MODEL 2: RANCH HANDS - INITIAL DIOXIN - UNADJUSTED

Initial Dioxin Category Summary Statistics		Analysis Results for Log ₂ (Initial Dioxin) ^a	
Initial Dioxin	n	Number (%) Abnormal	Estimated Relative Risk (95% C.I.) ^b	p-Value
Low	155	12 (7.7)	0.90 (0.69,1.17)	0.417
Medium	161	16 (9.9)		
High	160	12 (7.5)		

^a Adjusted for percent body fat at the time of the blood measurement of dioxin. ^b Relative risk for a twofold increase in initial dioxin.

Note: Low = 27-63 ppt; Medium = >63-152 ppt; High = >152 ppt.

(d) MODEL 2: RANCH HANDS – INITIAL DIOXIN – ADJUSTED

Analysis Results for Log ₂ (Initial Dioxin)						
	Adjusted Relative Risk					
n	(95% C.I.) ^a	p-Value				
457	1.11 (0.78,1.57)	0.561				

^a Relative risk for a twofold increase in initial dioxin.

Table 14-26. Analysis of Dorsalis Pedis Pulses (Continued)

(e) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY - UNADJUSTED

		Number (%)	Est. Relative Risk	
Dioxin Category	n	Abnormal	(95% C.I.) ^{ab}	p-Value
Comparison	1,193	95 (8.0)		
Background RH	376	29 (7.7)	0.91 (0.59,1.40)	0.664
Low RH	233	22 (9.4)	1.22 (0.75,1.98)	0.429
High RH	243	18 (7.4)	0.98 (0.58,1.65)	0.931
Low plus High RH	476	40 (8.4)	1.09 (0.74,1.61)	0.670

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: 1987 Dioxin ≤ 10 ppt.

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, $10 \text{ ppt} < \text{Initial Dioxin} \le 94 \text{ ppt}$.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

(f) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY - ADJUSTED

		Adjusted Relative Risk	
Dioxin Category	n	" (95% C.I.) ^a	p-Value
Comparison	1,153		
Background RH	360	0.94 (0.59,1.50)	0.792
Low RH	221	0.99 (0.58,1.70)	0.977
High RH	236	0.89 (0.50,1.58)	0.685
Low plus High RH	457	0.94 (0.61,1.43)	0.761

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: $1987 \text{ Dioxin} \le 10 \text{ ppt.}$

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, $10 \text{ ppt} < \text{Initial Dioxin} \le 94 \text{ ppt}$.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

(g) MODEL 4: RANCH HANDS – 1987 DIOXIN – UNADJUSTED

1987 Dioxin Category Summary Statistics		Analysis Results for Log ₂	(1987 Dioxin + 1)	
1987 Dioxin	n	Number (%) Abnormal	Estimated Relative Risk (95% C.I.) ^a	p-Value
Low	284	21 (7.4)	0.99 (0.84,1.17)	0.913
Medium	281	25 (8.9)		
High	287	23 (8.0)		

^a Relative risk for a twofold increase in 1987 dioxin.

Note: Low = \le 7.9 ppt; Medium = >7.9–19.6 ppt; High = >19.6 ppt.

^b Adjusted for percent body fat at the time of the blood measurement of dioxin.

Table 14-26. Analysis of Dorsalis Pedis Pulses (Continued)

(h) MODEL 4: RANCH HANDS – 1987 DIOXIN – ADJUSTED					
	Analysis Results for Log ₂ (1987 Dioxin + 1)				
	Adjusted Relative Risk				
n	(95% C.I.) ^a	p-Value			
817	1.07 (0.85,1.33)	0.580			

^a Relative risk for a twofold increase in 1987 dioxin.

14.2.2.3.7 Posterior Tibial Pulses

All unadjusted and adjusted Models 1 through 4 analyses of posterior tibial pulses were nonsignificant (Table 14-27(a–h): p>0.11 for each analysis).

Table 14-27. Analysis of Posterior Tibial Pulses

(a) MODEL 1: RANCH HANDS VS. COMPARISONS – UNADJUSTED					
Occupational Category	Group	n	Number (%) Abnormal	Est. Relative Risk (95% C.I.)	p-Value
All	Ranch Hand Comparison	859 1,228	58 (6.8) 64 (5.2)	1.32 (0.91,1.90)	0.142
Officer	Ranch Hand Comparison	334 483	22 (6.6) 23 (4.8)	1.41 (0.77,2.57)	0.263
Enlisted Flyer	Ranch Hand Comparison	149 183	14 (9.4) 13 (7.1)	1.36 (0.62,2.98)	0.449
Enlisted Groundcrew	Ranch Hand Comparison	376 562	22 (5.9) 28 (5.0)	1.19 (0.67,2.10)	0.562

(b) MODEL 1: RANCH HANDS VS. COMPARISONS – ADJUSTED					
Adjusted Relative Risk Occupational Category (95% C.I.) p-Value					
All	1.25 (0.84,1.86)	0.280			
Officer	1.40 (0.73,2.68)	0.307			
Enlisted Flyer	1.17 (0.49,2.78)	0.724			
Enlisted Groundcrew	1.16 (0.62,2.16)	0.649			

(c) MODEL 2:	(c) MODEL 2: RANCH HANDS – INITIAL DIOXIN – UNADJUSTED					
Initial Dioxin Category Summary Statistics			Analysis Results for Log ₂ (Initial Dioxin) ^a		
Initial Dioxin	n	Number (%) Abnormal	Estimated Relative Risk (95% C.I.) ^b	p-Value		
Low	155	9 (5.8)	1.01 (0.77,1.33)	0.925		
Medium	161	15 (9.3)				
High	160	10 (6.3)				

^a Adjusted for percent body fat at the time of the blood measurement of dioxin.

Note: Low = 27-63 ppt; Medium = >63-152 ppt; High = >152 ppt.

^b Relative risk for a twofold increase in initial dioxin.

Table 14-27. Analysis of Posterior Tibial Pulses (Continued)

(d) MODEL 2: RANCH HANDS – INITIAL DIOXIN – ADJUSTED				
	Analysis Results for Log_2 (Initial Diox	in)		
	Adjusted Relative Risk			
n	(95% C.I.) ^a	p-Value		
457	1.16 (0.81.1.65)	0.417		

^a Relative risk for a twofold increase in initial dioxin.

(e) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY – UNADJUSTED

Dioxin Category	n	Number (%) Abnormal	Est. Relative Risk (95% C.I.) ^{ab}	p-Value
Comparison	1,191	63 (5.3)		
Background RH	376	22 (5.9)	1.04 (0.63,1.73)	0.865
Low RH	233	18 (7.7)	1.52 (0.88, 2.61)	0.135
High RH	243	16 (6.6)	1.34 (0.76,2.36)	0.320
Low plus High RH	476	34 (7.1)	1.42 (0.92,2.19)	0.113

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: $1987 \text{ Dioxin} \le 10 \text{ ppt.}$

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, $10 \text{ ppt} < \text{Initial Dioxin} \le 94 \text{ ppt}$.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

(f) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY - ADJUSTED

	Adjusted Relative Risk		
Dioxin Category	n	(95% C.I.) ^a	p-Value
Comparison	1,151		
Background RH	360	1.08 (0.62,1.89)	0.784
Low RH	221	1.31 (0.71,2.39)	0.387
High RH	236	1.21 (0.63,2.30)	0.571
Low plus High RH	457	1.25 (0.77,2.03)	0.358

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: $1987 \text{ Dioxin} \le 10 \text{ ppt.}$

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, 10 ppt < Initial Dioxin ≤ 94 ppt.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

^b Adjusted for percent body fat at the time of the blood measurement of dioxin.

Table 14-27. Analysis of Posterior Tibial Pulses (Continued)

(g) MODEL 4: RANCH HANDS – 1987 DIOXIN – UNADJUSTED					
1987 Dioxin Category Summary Statistics Analysis Results for Log ₂ (1987				(1987 Dioxin + 1)	
1987 Dioxin	n	Number (%) Abnormal	Estimated Relative Risk (95% C.I.) ^a	p-Value	
Low	284	18 (6.3)	1.03 (0.86,1.24)	0.746	
Medium	281	16 (5.7)			
High	287	22 (7.7)			

^a Relative risk for a twofold increase in 1987 dioxin.

Note: Low = ≤ 7.9 ppt; Medium = > 7.9 - 19.6 ppt; High = > 19.6 ppt.

(h) MODEL 4: RANCH HANDS – 1987 DIOXIN – ADJUSTED				
	Analysis Results for Log ₂ (1987 Dioxin + 1)			
	Adjusted Relative Risk			
n	(95% C.I.) ^a	p-Value		
817	1.12 (0.88,1.43)	0.354		

^a Relative risk for a twofold increase in 1987 dioxin.

14.2.2.3.8 Leg Pulses

Leg pulses were not significantly associated with dioxin in any of the unadjusted and adjusted Models 1 through 4 analyses (Table 14-28(a–h): p>0.15 for each analysis).

Table 14-28. Analysis of Leg Pulses

(a) MODEL 1: RANCH HANDS VS. COMPARISONS – UNADJUSTED					
Occupational Category	Group	n	Number (%) Abnormal	Est. Relative Risk (95% C.I.)	p-Value
All	Ranch Hand Comparison	859 1,228	94 (10.9) 123 (10.0)	1.10 (0.83,1.47)	0.496
Officer	Ranch Hand Comparison	334 483	36 (10.8) 40 (8.3)	1.34 (0.83,2.15)	0.228
Enlisted Flyer	Ranch Hand Comparison	149 183	25 (16.8) 22 (12.0)	1.48 (0.79,2.74)	0.218
Enlisted Groundcrew	Ranch Hand Comparison	376 562	33 (8.8) 61 (10.9)	0.79 (0.51,1.23)	0.300

Table 14-28. Analysis of Leg Pulses (Continued)

(b) MODEL 1: RANCH HANDS VS. COMPARISONS - ADJUSTED

	Adjusted Relative Risk	
Occupational Category	(95% C.I.)	p-Value
All	1.03 (0.76,1.40)	0.850
Officer	1.30 (0.79,2.16)	0.306
Enlisted Flyer	1.46 (0.74,2.88)	0.270
Enlisted Groundcrew	0.71 (0.44,1.14)	0.158

(c) MODEL 2: RANCH HANDS – INITIAL DIOXIN – UNADJUSTED

Initial Dioxin Category Summary Statistics			Analysis Results for Log ₂ (Initial Dioxin) ^a
Initial Dioxin	n	Number (%) Abnormal	Estimated Relative Risk (95% C.I.) ^b	p-Value
Low	155	15 (9.7)	0.96 (0.77,1.20)	0.739
Medium	161	22 (13.7)		
High	160	16 (10.0)		

^a Adjusted for percent body fat at the time of the blood measurement of dioxin. ^b Relative risk for a twofold increase in initial dioxin.

Note: Low = 27-63 ppt; Medium = >63-152 ppt; High = >152 ppt.

(d) MODEL 2: RANCH HANDS - INITIAL DIOXIN - ADJUSTED

` '		
	Analysis Results for Log ₂ (Initial Diox	in)
	Adjusted Relative Risk	
n	(95% C.I.) ^a	p-Value
457	1.13 (0.84,1.51)	0.433

^a Relative risk for a twofold increase in initial dioxin.

(e) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY - UNADJUSTED

		Number (%)	Est. Relative Risk	
Dioxin Category	n	Abnormal	(95% C.I.) ^{ab}	p-Value
Comparison	1,191	122 (10.2)		
Background RH	376	39 (10.4)	0.95 (0.65,1.40)	0.812
Low RH	233	29 (12.4)	1.26 (0.82,1.94)	0.298
High RH	243	24 (9.9)	1.01 (0.64,1.61)	0.957
Low plus High RH	476	53 (11.1)	1.13 (0.80,1.59)	0.498

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: $1987 \text{ Dioxin} \le 10 \text{ ppt.}$

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, $10 \text{ ppt} < \text{Initial Dioxin} \le 94 \text{ ppt}$.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

^b Adjusted for percent body fat at the time of the blood measurement of dioxin.

Table 14-28. Analysis of Leg Pulses (Continued)

(f) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY - ADJUSTED

		Adjusted Relative Risk	
Dioxin Category	n	" (95% C.I.) ^a	p-Value
Comparison	1,151		
Background RH	360	1.01 (0.66,1.53)	0.981
Low RH	221	1.01 (0.63,1.64)	0.955
High RH	236	0.91 (0.54,1.53)	0.725
Low plus High RH	457	0.96 (0.66,1.40)	0.832

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: 1987 Dioxin ≤ 10 ppt.

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, 10 ppt < Initial Dioxin ≤ 94 ppt. High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

(g) MODEL 4: RANCH HANDS – 1987 DIOXIN – UNADJUSTED

(g) MODEL 4.		1507 1507	CIVIDGOSTED	
1987 Dioxin Category Summary Statistics		Analysis Results for Log ₂	(1987 Dioxin + 1)	
1987 Dioxin	n	Number (%) Abnormal	Estimated Relative Risk (95% C.I.) ^a	p-Value
Low	284	30 (10.6)	1.00 (0.87,1.16)	0.956
Medium	281	31 (11.0)		
High	287	31 (10.8)		

^a Relative risk for a twofold increase in 1987 dioxin.

Note: Low = \le 7.9 ppt; Medium = >7.9–19.6 ppt; High = >19.6 ppt.

(h) MODEL 4: RANCH HANDS – 1987 DIOXIN – ADJUSTED				
	Analysis Results for Log ₂ (1987 Dioxin + 1)			
	Adjusted Relative Risk			
n	(95% C.I.) ^a	p-Value		
817	1.08 (0.88,1.31)	0.467		

^a Relative risk for a twofold increase in 1987 dioxin.

14.2.2.3.9 Peripheral Pulses

All unadjusted and adjusted analyses in Models 1 through 4 were nonsignificant (Table 14–29(a–h): p>0.21 for each analysis).

Table 14-29. Analysis of Peripheral Pulses

(a) MODEL 1:	RANCH HANDS VS.	COMPARISONS -	UNADJUSTED

Occupational Category	Group	n	Number (%) Abnormal	Est. Relative Risk (95% C.I.)	p-Value
All	Ranch Hand Comparison	859 1,228	97 (11.3) 126 (10.3)	1.11 (0.84,1.47)	0.454
Officer	Ranch Hand Comparison	334 483	37 (11.1) 42 (8.7)	1.31 (0.82,2.08)	0.258
Enlisted Flyer	Ranch Hand Comparison	149 183	25 (16.8) 22 (12.0)	1.48 (0.79,2.74)	0.218
Enlisted Groundcrew	Ranch Hand Comparison	376 562	35 (9.3) 62 (11.0)	0.83 (0.53,1.28)	0.396

(b) MODEL 1: RANCH HANDS VS. COMPARISONS – ADJUSTED

	Adjusted Relative Risk	
Occupational Category	(95% C.I.)	p-Value
All	1.05 (0.77,1.42)	0.761
Officer	1.27 (0.77,2.09)	0.353
Enlisted Flyer	1.48 (0.75,2.92)	0.260
Enlisted Groundcrew	0.75 (0.47,1.21)	0.242

(c) MODEL 2: RANCH HANDS – INITIAL DIOXIN – UNADJUSTED

Initial Dioxin Category Summary Statistics		Analysis Results for Log ₂ (Initial Dioxin) ^a		
Initial Dioxin	n	Number (%) Abnormal	Estimated Relative Risk (95% C.I.) ^b	p-Value
Low	155	16 (10.3)	0.96 (0.77,1.19)	0.703
Medium	161	22 (13.7)		
High	160	17 (10.6)		

^a Adjusted for percent body fat at the time of the blood measurement of dioxin. ^b Relative risk for a twofold increase in initial dioxin.

Note: Low = 27-63 ppt; Medium = >63-152 ppt; High = >152 ppt.

(d) MODEL 2: RANCH HANDS – INITIAL DIOXIN – ADJUSTED Analysis Results for Log₂ (Initial Dioxin)

n	Adjusted Relative Risk (95% C.I.) ^a	p-Value
457	1.06 (0.79,1.41)	0.718

^a Relative risk for a twofold increase in initial dioxin.

Table 14-29. Analysis of Peripheral Pulses (Continued)

(e) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY - UNADJUSTED

		Number (%)	Est. Relative Risk	
Dioxin Category	n	Abnormal	(95% C.I.) ^{ab}	p-Value
Comparison	1,191	125 (10.5)		
Background RH	376	40 (10.6)	0.95 (0.65,1.39)	0.797
Low RH	233	30 (12.9)	1.27 (0.83,1.95)	0.266
High RH	243	25 (10.3)	1.04 (0.66,1.63)	0.880
Low plus High RH	476	55 (11.6)	1.15 (0.82,1.61)	0.431

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: 1987 Dioxin ≤ 10 ppt.

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, 10 ppt < Initial Dioxin ≤ 94 ppt.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

(f) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY - ADJUSTED

		Adjusted Relative Risk	
Dioxin Category	n	(95% C.I.) ^a	p-Value
Comparison	1,151		
Background RH	360	1.00 (0.66,1.52)	0.997
Low RH	221	1.05 (0.65,1.70)	0.833
High RH	236	0.94 (0.57,1.57)	0.828
Low plus High RH	457	1.00 (0.68,1.45)	0.981

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: 1987 Dioxin ≤ 10 ppt.

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, 10 ppt < Initial Dioxin ≤ 94 ppt.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

(g) MODEL 4: RANCH HANDS – 1987 DIOXIN – UNADJUSTED

1987 Dioxin Category Summary Statistics		Analysis Results for Log ₂	(1987 Dioxin + 1)	
1987 Dioxin	n	Number (%) Abnormal	Estimated Relative Risk (95% C.I.) ^a	p-Value
Low	284	31 (10.9)	1.00 (0.86,1.15)	0.972
Medium	281	32 (11.4)		
High	287	32 (11.1)		

^a Relative risk for a twofold increase in 1987 dioxin.

Note: Low = ≤ 7.9 ppt; Medium = > 7.9 - 19.6 ppt; High = > 19.6 ppt.

^b Adjusted for percent body fat at the time of the blood measurement of dioxin.

Table 14-29. Analysis of Peripheral Pulses (Continued)

(h) MODEL 4: RANCH HANDS – 1987 DIOXIN – ADJUSTED						
	Analysis Results for Log_2 (1987 Dioxin + 1)					
	Adjusted Relative Risk					
n	(95% C.I.) ^a	p-Value				
817	1.07 (0.88,1.30)	0.485				

^a Relative risk for a twofold increase in 1987 dioxin.

14.2.2.3.10 ICVI Index

The analysis of ICVI index did not show any significant associations with dioxin (Table 14-30(a-h): p>0.11 for each analysis).

Table 14-30. Analysis of ICVI Index

(a) MODEL 1:	(a) MODEL 1: RANCH HANDS VS. COMPARISONS – UNADJUSTED					
Occupational Category	Group	n	Number (%) Abnormal	Est. Relative Risk (95% C.I.)	p-Value	
All	Ranch Hand Comparison	858 1,232	33 (3.8) 45 (3.7)	1.06 (0.67,1.67)	0.819	
Officer	Ranch Hand Comparison	334 484	13 (3.9) 15 (3.1)	1.27 (0.59,2.70)	0.541	
Enlisted Flyer	Ranch Hand Comparison	149 186	7 (4.7) 12 (6.5)	0.71 (0.27,1.86)	0.492	
Enlisted Groundcrew	Ranch Hand Comparison	375 562	13 (3.5) 18 (3.2)	1.09 (0.53,2.24)	0.825	

(b) MODEL 1: RANCH HANDS VS. COMPARISONS – ADJUSTED					
Occupational Category	Adjusted Relative Risk (95% C.I.)	p-Value			
All	0.99 (0.61,1.60)	0.958			
Officer	1.25 (0.57,2.70)	0.577			
Enlisted Flyer	0.50 (0.17,1.51)	0.218			
Enlisted Groundcrew	1.12 (0.53,2.39)	0.764			

(c) MODEL 2: RANCH HANDS – INITIAL DIOXIN – UNADJUSTED					
Initial Dioxin Category Summary Statistics Analysis Results for Log ₂ (Initial Dioxin) ^a					
Initial Dioxin	n	Number (%) Abnormal	Estimated Relative Risk (95% C.I.) ^b	p-Value	
Low	155	6 (3.9)	0.99 (0.71,1.37)	0.948	
Medium	161	10 (6.2)			
High	160	7 (4.4)			

^a Adjusted for percent body fat at the time of the blood measurement of dioxin. ^b Relative risk for a twofold increase in initial dioxin.

Note: Low = 27-63 ppt; Medium = >63-152 ppt; High = >152 ppt.

Table 14-30. Analysis of ICVI Index (Continued)

(d) MODEL 2: RANCH HANDS – INITIAL DIOXIN – ADJUSTED							
	Analysis Results for Log ₂ (Initial Diox	in)					
n	Adjusted Relative Risk	n-Valua					

0.604

461

Note: Results are not adjusted for diabetic class because of the sparse number of Ranch Hands with an abnormal intermittent claudication and vascular insufficiency index.

1.12 (0.73,1.72)

(e) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY - UNADJUSTED

		Number (%)	Est. Relative Risk	
Dioxin Category	n	Abnormal	(95% C.I.) ^{ab}	p-Value
Comparison	1,195	43 (3.6)		
Background RH	375	9 (2.4)	0.65 (0.31,1.35)	0.249
Low RH	233	9 (3.9)	1.08 (0.52,2.24)	0.839
High RH	243	14 (5.8)	1.66 (0.89,3.09)	0.112
Low plus High RH	476	23 (4.8)	1.34 (0.79,2.27)	0.272

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: 1987 Dioxin ≤ 10 ppt.

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, 10 ppt < Initial Dioxin ≤ 94 ppt.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

(f) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY - ADJUSTED

		Adjusted Relative Risk	
Dioxin Category	n	(95% C.I.) ^a	p-Value
Comparison	1,155		
Background RH	360	0.69 (0.32,1.48)	0.340
Low RH	221	0.98 (0.46,2.11)	0.968
High RH	236	1.41 (0.69,2.89)	0.346
Low plus High RH	457	1.19 (0.67,2.09)	0.555

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: $1987 \text{ Dioxin} \le 10 \text{ ppt.}$

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, 10 ppt < Initial Dioxin ≤ 94 ppt.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

^a Relative risk for a twofold increase in initial dioxin.

^b Adjusted for percent body fat at the time of the blood measurement of dioxin.

Table 14-30. Analysis of ICVI Index (Continued)

(g) MODEL 4: RANCH HANDS – 1987 DIOXIN – UNADJUSTED								
1987 Dioxir	n Category Sum	mary Statistics	Analysis Results for Log ₂ (1987 Dioxin + 1)					
1987 Dioxin	n	Number (%) Abnormal	Estimated Relative Risk (95% C.I.) ^a	p-Value				
Low	283	8 (2.8)	1.08 (0.86,1.37)	0.503				
Medium	281	9 (3.2)						

^a Relative risk for a twofold increase in 1987 dioxin.

287

High

Note: Low = ≤ 7.9 ppt; Medium = > 7.9 - 19.6 ppt; High = > 19.6 ppt.

15 (5.2)

(h) MODEL 4: RANCH HANDS – 1987 DIOXIN – ADJUSTED							
Analysis Results for Log ₂ (1987 Dioxin + 1)							
Adjusted Relative Risk							
n	(95% C.I.) ^a	p-Value					
817	1.07 (0.79,1.45)	0.666					

^a Relative risk for a twofold increase in 1987 dioxin.

14.2.3 Longitudinal Analysis

Cardiovascular longitudinal analyses were conducted on systolic blood pressure measurements taken at the 1982 and 1997 examinations and six pulse assessments made at the 1985 and 1997 examinations. Discrete and continuous analyses were performed for systolic blood pressure. The six pulse measurements included femoral pulses, popliteal pulses, dorsalis pedis pulses, posterior tibial pulses, leg pulses, and peripheral pulses. The 1985 and 1997 measurements were used for the pulse assessments because the Doppler assessment of pulses was conducted at these two examinations and was not conducted at the 1982 baseline or 1987 follow-up examinations.

Longitudinal analyses were conducted to examine whether changes across time differed with respect to group membership (Model 1), initial dioxin (Model 2), and categorized dioxin (Model 3). Model 4 was not examined in longitudinal analyses because 1987 dioxin, the measure of exposure in these models, changes over time and is not available for all participants for 1982 or 1997.

Participants considered abnormal in 1982 (or 1985 for Doppler pulse measurements) were not included in the longitudinal analysis of discrete dependent variables. The purpose of the longitudinal analysis was to examine the effects of dioxin exposure across time. Participants who were abnormal in 1982 (or 1988) were not considered to be at risk for developing the condition, because the condition already existed at the time of the first collection of data for the AFHS (1982). Only participants who were normal at the 1982 (or 1985) examination were considered to be at risk for developing the condition; therefore, the rate of abnormalities under this restriction approximates an incidence rate between 1982 (or 1985) and 1997. That is, an incidence rate is a measure of the rate at which people without a condition develop the condition during a specified period of time (53). Summary statistics are provided for reference purposes for the 1985, 1987, and 1992 examinations for systolic blood pressure and for the 1992 examination for the pulse measurements.

The longitudinal analysis for systolic blood pressure in its discrete form examined relative risks at the 1997 examination for participants who were classified as normal at the 1982 examination. The longitudinal analysis for the Doppler pulse measurements examined relative risks at the 1997 examination for participants who were classified as normal at the 1985 examination. The adjusted relative risks estimated from each of the three models were used to investigate the change in the dependent variable over time. All three models were adjusted for age; Models 2 and 3 also were adjusted for the percentage of body fat at the time of the blood measurement of dioxin.

The longitudinal analysis for the systolic blood pressure in its continuous form examined the paired difference between the measurements from 1982 and 1997. These paired differences measured the change in systolic blood pressure over time. Each of the three models used in the longitudinal analysis was adjusted for age and systolic blood pressure as measured in 1982 (see Chapter 7, Statistical Methods).

14.2.3.1 Physical Examination Variables

14.2.3.1.1 Systolic Blood Pressure (Continuous)

The Model 1 analysis of change in mean systolic blood pressure revealed a marginally significant difference between overall Ranch Hands and Comparisons (Table 14-31(a): difference of examination mean change=–1.6 mm Hg, p=0.066). The Ranch Hand mean decreased by 6.3 mm Hg between 1982 and 1997, and the Comparison mean decreased by 4.7 mm Hg. Stratifying by occupation showed a marginally significant group difference in the enlisted groundcrew stratum (Table 14-31(a): difference of examination mean change=–2.2 mm Hg, p=0.079). For the enlisted groundcrew, the Ranch Hand mean decreased by 7.4 mm Hg between 1982 and 1997, and the Comparison mean decreased by 5.2 mm Hg.

Table 14-31. Longitudinal Analysis of Systolic Blood Pressure (mm Hg) (Continuous)

Occupational				Mean ^a /(n) xaminatio		Exam. Mean	Difference of Exam. Mean		
Category	Group	1982	1985	1987	1992	1997	Change ^b	Change	p-Value
All	Ranch Hand	131.1	117.8	125.9	120.4	124.8	-6.3	-1.6	0.066
	Comparison	(808) 130.7 (959)	(790) 118.9 (940)	(782) 126.4 (935)	(785) 121.3 (939)	(808) 126.0 (959)	<i>−4.7</i>		
Officer	Ranch Hand	131.8 (305)	118.8 (301)	126.5 (298)	122.6 (300)	126.1 (305)	-5.6	-0.3	0.840
	Comparison	131.3 (372)	118.8 (365)	126.3 (360)	121.8 (367)	126.1 (372)	-5.3		
Enlisted Flyer	Ranch Hand	131.8 (146)	118.4 (143)	127.2 (141)	120.6 (142)	126.7 (146)	-5.1	-3.8	0.135
	Comparison	130.2 (144)	118.9 (143)	125.9 (142)	121.2 (142)	128.9 (144)	-1.3		
Enlisted Groundcrew	Ranch Hand	130.3 (357)	116.8 (346)	124.8 (343)	118.4 (343)	122.9 (357)	-7.4	-2.2	0.079
	Comparison	130.3 (443)	119.0 (432)	126.7 (433)	120.9 (430)	125.1 (443)	-5.2		

Note: Summary statistics for 1985 are provided for reference purposes for participants who attended the 1982, 1985, and 1997 examinations. Summary statistics for 1987 are provided for reference purposes for participants who attended the 1982, 1987, and 1997 examinations. Summary statistics for 1992 are provided for reference purposes for participants who attended the 1982, 1992, and 1997 examinations.

 ^a Transformed from natural logarithm scale.
 ^b Difference between 1997 and 1982 examination means after transformation to original scale.

^c P-value is based on analysis of natural logarithm of systolic blood pressure; results adjusted for natural logarithm of systolic blood pressure in 1982 and age in 1997.

Table 14-31. Longitudinal Analysis of Systolic Blood Pressure (mm Hg) (Continuous) (Continued)

(b) MODEL 2	(b) MODEL 2: RANCH HANDS – INITIAL DIOXIN										
Ini	Initial Dioxin Category Summary Statistics					Analysis Results for Log ₂ (I	(nitial Dioxin) ^b				
		I	Mean ^a /(n) Examinatio	n		Adjusted Slope					
Initial Dioxin	1982	1985	1987	1992	1997	(Std. Error)	p-Value				
Low	132.2 (149)	118.4 (146)	127.1 (148)	120.5 (144)	125.9 (149)	0.000 (0.005)	0.977				
Medium	132.8 (158)	119.7 (155)	126.4 (155)	122.9 (155)	125.5 (158)						
High	131.2 (153)	119.1 (150)	127.4 (148)	121.1 (150)	124.1 (153)						

^a Transformed from natural logarithm scale.

Note: Low = 27-63 ppt; Medium = >63-152 ppt; High = >152 ppt.

Summary statistics for 1985 are provided for reference purposes for participants who attended the 1982, 1985, and 1997 examinations. Summary statistics for 1987 are provided for reference purposes for participants who attended the 1982, 1987, and 1997 examinations. Summary statistics for 1992 are provided for reference purposes for participants who attended the 1982, 1992, and 1997 examinations.

^b Results based on difference between natural logarithm of 1997 systolic blood pressure and natural logarithm of 1982 systolic blood pressure versus log₂ (initial dioxin); results adjusted for percent body fat at the date of the blood measurement of dioxin, natural logarithm of 1982 systolic blood pressure, and age in 1997.

Table 14-31. Longitudinal Analysis of Systolic Blood Pressure (mm Hg) (Continuous) (Continued)

(c) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY

Dioxin -		I	Mean ^a /(n) Examination	n	Exam. Mean	Difference of Exam. Mean		
Category	1982	1985	1987	1992	1997	Change ^b	Change	p-Value ^c
Comparison	130.6	118.7	126.2	121.1	126.0	-4.7		
	(932)	(916)	(910)	(913)	(932)			
Background RH	129.8	116.2	124.4	119.0	124.4	-5.3	-0.6	0.386
	(342)	(334)	(326)	(331)	(342)			
Low RH	132.0	118.7	126.8	120.9	126.0	-6.0	-1.3	0.347
	(224)	(218)	(221)	(217)	(224)			
High RH	132.1	119.5	127.2	122.0	124.4	-7.8	-3.1	0.086
	(236)	(233)	(230)	(232)	(236)			
Low plus High RH	132.1	119.1	127.0	121.5	125.2	-6.9	-2.2	0.083
	(460)	(451)	(451)	(449)	(460)			

^a Transformed from natural logarithm scale.

Note: RH = Ranch Hand.

Comparison: $1987 \text{ Dioxin} \le 10 \text{ ppt.}$

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, 10 ppt < Initial Dioxin ≤ 94 ppt.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

Summary statistics for 1985 are provided for reference purposes for participants who attended the 1982, 1985, and 1997 examinations. Summary statistics for 1987 are provided for reference purposes for participants who attended the 1982, 1987, and 1997 examinations. Summary statistics for 1992 are provided for reference purposes for participants who attended the 1982, 1992, and 1997 examinations.

The longitudinal analysis in Model 2 did not reveal a significant association between the change in mean systolic blood pressure and dioxin (Table 14-31(b): p=0.977).

The Model 3 analysis of the change in mean systolic blood pressure levels between 1982 and 1997 revealed two marginally significant contrasts: Ranch Hands in the high dioxin category versus Comparisons (Table 14-31(c): difference of examination mean change=–3.1 mm Hg, p=0.086) and Ranch Hands in the low plus high dioxin category versus Comparisons (Table 14-31(c): difference of examination mean change=–2.2 mm Hg, p=0.083). The change in means between 1982 and 1997 for Ranch Hands in the high dioxin category, Ranch Hands in the low plus high dioxin category, and Comparisons was –7.8 mm Hg, –6.9 mm Hg, and –4.7 mm Hg, respectively.

14.2.3.1.2 Systolic Blood Pressure (Discrete)

The longitudinal analysis in Models 1 through 3 did not reveal a significant association between dioxin and change in systolic blood pressure in its discrete form (Table 14-32(a–c): p>0.45 for each analysis).

^b Difference between 1997 and 1982 examination means after transformation to original scale.

^c P-value is based on analysis of natural logarithm of 1997 systolic blood pressure; results adjusted for percent body fat at the date of the blood measurement of dioxin, natural logarithm of 1982 systolic blood pressure, and age in 1997.

Table 14-32. Longitudinal Analysis of Systolic Blood Pressure (Discrete)

(a) MODEL 1: RANCH HANDS VS. COMPARISONS

Occupational		Number (%) High/(n) Examination						
Category	Group	1982	1985	1987	1992	1997		
All	Ranch Hand	141 (17.5) (808)	42 (5.3) (790)	146 (18.7) (782)	119 (15.2) (785)	169 (20.9) (808)		
	Comparison	187 (19.5) (959)	65 (6.9) (940)	205 (21.9) (935)	146 (15.5) (939)	215 (22.4) (959)		
Officer	Ranch Hand	60 (19.7) (305)	20 (6.6) (301)	59 (19.8) (298)	51 (17.0) (300)	73 (23.9) (305)		
	Comparison	75 (20.2) (372)	25 (6.8) (365)	81 (22.5) (360)	65 (17.7) (367)	90 (24.2) (372)		
Enlisted Flyer	Ranch Hand	28 (19.2) (146)	5 (3.5) (143)	29 (20.6) (141)	23 (16.2) (142)	35 (24.0) (146)		
	Comparison	27 (18.8) (144)	11 (7.7) (143)	31 (21.8) (142)	20 (14.1) (142)	38 (26.4) (144)		
Enlisted Groundcrew	Ranch Hand	53 (14.8) (357)	17 (4.9) (346)	58 (16.9) (343)	45 (13.1) (343)	61 (17.1) (357)		
	Comparison	85 (19.2) (443)	29 (6.7) (432)	93 (21.5) (433)	61 (14.2) (430)	87 (19.6) (443)		

	_	Nor	mal in 1982		
Occupational Category	Group	n in 1997	Number (%) High in 1997	Adj. Relative Risk (95% C.I.) ^a	p-Value ^a
All	Ranch Hand Comparison	667 772	111 (16.6) 130 (16.8)	0.99 (0.75,1.31)	0.951
Officer	Ranch Hand Comparison	245 297	48 (19.6) 50 (16.8)	1.18 (0.76,1.84)	0.454
Enlisted Flyer	Ranch Hand Comparison	118 117	23 (19.5) 25 (21.4)	0.90 (0.47,1.71)	0.743
Enlisted Groundcrew	Ranch Hand Comparison	304 358	40 (13.2) 55 (15.4)	0.86 (0.55,1.35)	0.513

^a Relative risk, confidence interval, and p-values are in reference to a contrast of 1982 and 1997 results; results adjusted for age in 1997.

Note: Summary statistics for 1985 are provided for reference purposes for participants who attended the 1982, 1985, and 1997 examinations. Summary statistics for 1987 are provided for reference purposes for participants who attended the 1982, 1987, and 1997 examinations. Summary statistics for 1992 are provided for reference purposes for participants who attended the 1982, 1992, and 1997 examinations. Statistical analyses are based only on participants who had normal systolic blood pressure in 1982 (see Chapter 7, Statistical Methods).

Table 14-32. Longitudinal Analysis of Systolic Blood Pressure (Discrete) (Continued)

(b) MODEL 2: RANCH HANDS — INIT	TAT	DIOXIN
---------------------------------	-----	--------

	Number (%) High/(n) Examination							
Initial Dioxin	1982	1985	1987	1992	1997			
Low	32 (21.5)	6 (4.1)	33 (22.3)	24 (16.7)	37 (24.8)			
	(149)	(146)	(148)	(144)	(149)			
Medium	32 (20.3)	8 (5.2)	28 (18.1)	28 (18.1)	34 (21.5)			
	(158)	(155)	(155)	(155)	(158)			
High	22 (14.4)	11 (7.3)	30 (20.3)	25 (16.7)	28 (18.3)			
C	(153)	(150)	(148)	(150)	(153)			

Initial	Dioxin Category St	ımmary Statistics	Analysis Results for Log	₂ (Initial Dioxin) ^a
	No	rmal in 1982		
Initial		Number (%) High	Adj. Relative Risk	
Dioxin	n in 1997	in 1997	(95% C.I.) ^b	p-Value
Low	117	22 (18.8)	0.96 (0.78,1.19)	0.714
Medium	126	23 (18.3)		
High	131	20 (15.3)		

^a Adjusted for percent body fat at the time of the blood measurement of dioxin and age in 1997.

Note: Low = 27-63 ppt; Medium = >63-152 ppt; High = >152 ppt.

Summary statistics for 1985 are provided for reference purposes for participants who attended the 1982, 1985, and 1997 examinations. Summary statistics for 1987 are provided for reference purposes for participants who attended the 1982, 1987, and 1997 examinations. Summary statistics for 1992 are provided for reference purposes for participants who attended the 1982, 1992, and 1997 examinations. Statistical analyses are based only on participants who had normal systolic blood pressure in 1982 (see Chapter 7, Statistical Methods).

(c) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY

]	Number (%) High/(r Examination	1)	
Dioxin Category	1982	1985	1987	1992	1997
Comparison	180 (19.3)	60 (6.6)	194 (21.3)	140 (15.3)	207 (22.2)
	(932)	(916)	(910)	(913)	(932)
Background RH	54 (15.8)	17 (5.1)	54 (16.6)	42 (12.7)	69 (20.2)
•	(342)	(334)	(326)	(331)	(342)
Low RH	43 (19.2)	8 (3.7)	44 (19.9)	35 (16.1)	54 (24.1)
	(224)	(218)	(221)	(217)	(224)
High RH	43 (18.2)	17 (7.3)	47 (20.4)	42 (18.1)	45 (19.1)
•	(236)	(233)	(230)	(232)	(236)
Low plus High RH	86 (18.7)	25 (5.5)	91 (20.2)	77 (17.1)	99 (21.5)
	(460)	(451)	(451)	(449)	(460)

^b Relative risk for a twofold increase in initial dioxin.

Table 14-32. Longitudinal Analysis of Systolic Blood Pressure (Discrete) (Continued)

	Norma	l in 1982		
Dioxin Category	n in 1997	Number (%) High in 1997	Adj. Relative Risk (95% C.I.) ^{ab}	p-Value ^b
Comparison	752	127 (16.9)		
Background RH	288	45 (15.6)	0.96 (0.66,1.41)	0.840
Low RH	181	34 (18.8)	1.01 (0.65,1.55)	0.978
High RH	193	31 (16.1)	1.01 (0.65,1.57)	0.965
Low plus High RH	374	65 (17.4)	1.01 (0.72,1.41)	0.963

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: 1987 Dioxin ≤ 10 ppt.

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin >10 ppt, 10 ppt < Initial Dioxin ≤ 94 ppt.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

Summary statistics for 1985 are provided for reference purposes for participants who attended the 1982, 1985, and 1997 examinations. Summary statistics for 1987 are provided for reference purposes for participants who attended the 1982, 1987, and 1997 examinations. Summary statistics for 1992 are provided for reference purposes for participants who attended the 1982, 1992, and 1997 examinations. Statistical analyses are based only on participants who had normal systolic blood pressure in 1982 (see Chapter 7, Statistical Methods).

14.2.3.1.3 Femoral Pulses

The Model 1 analysis of the change in percentage of abnormal femoral pulses did not reveal a significant difference between Ranch Hands and Comparisons overall (Table 14-33(a): p=0.118). Stratifying by occupation showed a marginally significant group difference in the enlisted groundcrew stratum (Table 14-33(a): Adj. RR=3.19, p=0.095). For enlisted groundcrew, 1.9 percent of the Ranch Hands and 0.6 percent of the Comparisons had normal femoral pulses in 1985 and abnormal femoral pulses in 1997.

The Model 2 longitudinal analysis revealed no significant association between dioxin and the percentage of participants with normal femoral pulses in 1985 and abnormal femoral pulses in 1997 (Table 14-33(b): p=0.972).

^b Adjusted for percent body fat at the time of the blood measurement of dioxin and age in 1997

Table 14-33. Longitudinal Analysis of Femoral Pulses

	(a)	MODEL	1:	RANCH HANDS	VS. COMPARISONS
--	-----	-------	----	-------------	-----------------

Occupational]	Number (%) Abnormal/(n Examination	1)
Category	Group	1985	1992	1997
All	Ranch Hand	0 (0.0) (823)	6 (0.7) (802)	19 (2.3) (823)
	Comparison	0 (0.0) (1,047)	6 (0.6) (1,020)	14 (1.3) (1,047)
Officer	Ranch Hand	0 (0.0) (318)	4 (1.3) (313)	7 (2.2) (318)
	Comparison	0 (0.0) (412)	2 (0.5) (405)	8 (1.9) (412)
Enlisted Flyer	Ranch Hand	0 (0.0) (145)	0 (0.0) (143)	5 (3.4) (145)
	Comparison	0 (0.0) (158)	2 (1.3) (156)	3 (1.9) (158)
Enlisted Groundcrew	Ranch Hand	0 (0.0) (360)	2 (0.6) (346)	7 (1.9) (360)
	Comparison	0 (0.0) (477)	2 (0.4) (459)	3 (0.6) (477)

		Nor	mal in 1985		
Occupational Category	Group	n in 1997	Number (%) Abnormal in 1997	Adj. Relative Risk (95% C.I.) ^a	p-Value ^a
All	Ranch Hand Comparison	823 1,047	19 (2.3) 14 (1.3)	1.74 (0.86,3.49)	0.118
Officer	Ranch Hand Comparison	318 412	7 (2.2) 8 (1.9)	1.12 (0.40,3.13)	0.824
Enlisted Flyer	Ranch Hand Comparison	145 158	5 (3.4) 3 (1.9)	1.82 (0.43,7.77)	0.419
Enlisted Groundcrew	Ranch Hand Comparison	360 477	7 (1.9) 3 (0.6)	3.19 (0.82,12.42)	0.095

^a Relative risk, confidence interval, and p-values are in reference to a contrast of 1985 and 1997 results; results adjusted for age in 1997.

Note: Summary statistics for 1992 are provided for reference purposes for participants who attended the 1985 and 1997 examinations. Statistical analyses are based only on participants who had normal femoral pulses in 1985 (see Chapter 7, Statistical Methods).

Table 14-33. Longitudinal Analysis of Femoral Pulses (Continued)

(b)) MODEL 2:	RANCH HANDS -	— INITIAL DIOXIN
------------	------------	---------------	------------------

		Number (%) Abnormal/(n) Examination	
Initial Dioxin	1985	1992	1997
OW	0 (0.0)	3 (2.1)	3 (2.0)
	(149)	(144)	(149)
Iedium	0 (0.0)	1 (0.6)	5 (3.2)
	(158)	(155)	(158)
igh	0 (0.0)	0 (0.0)	4 (2.6)
	(155)	(151)	(155)

Initial	Dioxin Category Su	ımmary Statistics	Analysis Results for Log	₂ (Initial Dioxin) ^a
	No	rmal in 1985		
Initial Dioxin	n in 1997	Number (%) Abnormal in 1997	Adj. Relative Risk (95% C.I.) ^b	p-Value
Low	149	3 (2.0)	1.01 (0.63,1.61)	0.972
Medium	158	5 (3.2)		
High	155	4 (2.6)		

^a Adjusted for percent body fat at the time of the blood measurement of dioxin and age in 1997. ^b Relative risk for a twofold increase in initial dioxin.

Note: Low = 27-63 ppt; Medium = >63-152 ppt; High = >152 ppt.

Summary statistics for 1992 are provided for reference purposes for participants who attended the 1985 and 1997 examinations. Statistical analyses are based only on participants who had normal femoral pulses in 1985 (see Chapter 7, Statistical Methods).

(c) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY

		Number (%) Abnormal/(n) Examination	
Dioxin Category	1985	1992	1997
Comparison	0 (0.0)	6 (0.6)	14 (1.4)
_	(1,019)	(994)	(1,019)
Background RH	0 (0.0)	2 (0.6)	7 (2.0)
	(355)	(346)	(355)
Low RH	0 (0.0)	4 (1.8)	6 (2.7)
	(224)	(217)	(224)
High RH	0 (0.0)	0 (0.0)	6 (2.5)
	(238)	(233)	(238)
Low plus High RH	0 (0.0)	4 (0.9)	12 (2.6)
1 0	(462)	(450)	(462)

Table 14-33. Longitudinal Analysis of Femoral Pulses (Continued)

	Norm	nal in 1985		
Dioxin Category	n in 1997	Number (%) Abnormal in 1997	Adj. Relative Risk (95% C.I.) ^{ab}	p-Value ^b
Comparison	1,019	14 (1.4)		
Background RH	355	7 (2.0)	1.28 (0.51,3.21)	0.602
Low RH	224	6 (2.7)	1.88 (0.71,4.98)	0.202
High RH	238	6 (2.5)	2.34 (0.87,6.25)	0.091
Low plus High RH	462	12 (2.6)	2.10 (0.96,4.62)	0.063

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: $1987 \text{ Dioxin} \le 10 \text{ ppt.}$

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin >10 ppt, 10 ppt < Initial Dioxin ≤ 94 ppt.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

Summary statistics for 1992 are provided for reference purposes for participants who attended the 1985 and 1997 examinations. Statistical analyses are based only on participants who had normal femoral pulses in 1985 (see Chapter 7, Statistical Methods).

Model 3 analysis of the change in femoral pulses from normal in 1985 to abnormal in 1997 revealed two marginally significant contrasts: Ranch Hands in the high dioxin category versus Comparisons (Table 14-33(c): Adj. RR=2.34, p=0.091) and Ranch Hands in the low plus high dioxin category versus Comparisons (Table 14-33(c): Adj. RR=2.10, p=0.063). Of the Comparisons, 1.4 percent had normal femoral pulses in 1985 and abnormal femoral pulses in 1997. Of the Ranch Hands, 2.5 percent in the high dioxin category and 2.6 percent in the low plus high dioxin category had normal femoral pulses in 1985 and abnormal femoral pulses in 1997.

14.2.3.1.4 Popliteal Pulses

Analyses of Models 1 through 3 showed no significant associations between dioxin and the change in popliteal pulses between 1985 and 1997 (Table 14-34(a–c): p>0.19 for each analysis).

^b Adjusted for percent body fat at the time of the blood measurement of dioxin and age in 1997.

Table 14-34. Longitudinal Analysis of Popliteal Pulses

(a) MODEL 1: RANCH HANDS VS. COMPARISONS

Occupational		Number (%) Abnormal/(n) Examination				
Category	Group	1985	1992	1997		
All	Ranch Hand	2 (0.2) (823)	10 (1.2) (802)	23 (2.8) (823)		
	Comparison	1 (0.1) (1,046)	7 (0.7) (1,019)	24 (2.3) (1,046)		
Officer	Ranch Hand	1 (0.3) (318)	6 (1.9) (313)	7 (2.2) (318)		
	Comparison	0 (0.0) (411)	4 (1.0) (404)	11 (2.7) (411)		
Enlisted Flyer	Ranch Hand	0 (0.0) (145)	2 (1.4) (143)	5 (3.4) (145)		
	Comparison	1 (0.6) (158)	2 (1.3) (156)	3 (1.9) (158)		
Enlisted Groundcrew	Ranch Hand	1 (0.3) (360)	2 (0.6) (346)	11 (3.1) (360)		
	Comparison	0 (0.0) (477)	1 (0.2) (459)	10 (2.1) (477)		

	_	Normal in 1985			
Occupational Category	Group	n in 1997	Number (%) Abnormal in 1997	Adj. Relative Risk (95% C.I.) ^a	p-Value ^a
All	Ranch Hand Comparison	821 1,045	22 (2.7) 23 (2.2)	1.22 (0.67,2.21)	0.518
Officer	Ranch Hand Comparison	317 411	7 (2.2) 11 (2.7)	0.81 (0.31,2.13)	0.672
Enlisted Flyer	Ranch Hand Comparison	145 157	5 (3.4) 2 (1.3)	2.67 (0.51,14.07)	0.246
Enlisted Groundcrew	Ranch Hand Comparison	359 477	10 (2.8) 10 (2.1)	1.39 (0.57,3.40)	0.473

^a Relative risk, confidence interval, and p-values are in reference to a contrast of 1985 and 1997 results; results adjusted for age in 1997.

Note: Summary statistics for 1992 are provided for reference purposes for participants who attended the 1985 and 1997 examinations. Statistical analyses are based only on participants who had normal popliteal pulses in 1985 (see Chapter 7, Statistical Methods).

Table 14-34. Longitudinal Analysis of Popliteal Pulses (Continued)

(b) MODEL 2:	RANCH HAND	S — INITIAI	DIOXIN
--------------	------------	-------------	--------

		Number (%) Abnormal/(n) Examination	
Initial Dioxin	1985	1992	1997
Low	0 (0.0)	3 (2.1)	4 (2.7)
	(149)	(144)	(149)
Medium	0 (0.0)	2 (1.3)	6 (3.8)
	(158)	(155)	(158)
High	0 (0.0)	2(1.3)	4 (2.6)
	(155)	(151)	(155)

Initial	Dioxin Category Su	ımmary Statistics	Analysis Results for Log	₂ (Initial Dioxin) ^a
Normal in 1985				
Initial Dioxin	n in 1997	Number (%) Abnormal in 1997	Adj. Relative Risk (95% C.I.) ^b	p-Value
Low	149	4 (2.7)	0.95 (0.61,1.49)	0.838
Medium	158	6 (3.8)		
High	155	4 (2.6)		

^a Adjusted for percent body fat at the time of the blood measurement of dioxin and age in 1997. ^b Relative risk for a twofold increase in initial dioxin.

Note: Low = 27-63 ppt; Medium = >63-152 ppt; High = >152 ppt.

Summary statistics for 1992 are provided for reference purposes for participants who attended the 1985 and 1997 examinations. Statistical analyses are based only on participants who had normal popliteal pulses in 1985 (see Chapter 7, Statistical Methods).

(c) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY

		Number (%) Abnormal/(n) Examination	
Dioxin Category	1985	1992	1997
Comparison	1 (0.1)	7 (0.7)	24 (2.4)
•	(1,018)	(993)	(1,018)
Background RH	2 (0.6)	3 (0.9)	9 (2.5)
	(355)	(346)	(355)
Low RH	0 (0.0)	4 (1.8)	7 (3.1)
	(224)	(217)	(224)
High RH	0(0.0)	3 (1.3)	7 (2.9)
-	(238)	(233)	(238)
Low plus High RH	0(0.0)	7 (1.6)	14 (3.0)
1 3	(462)	(450)	(462)

Table 14-34. Longitudinal Analysis of Popliteal Pulses (Continued)

	Normal in 1985			
Dioxin Category	n in 1997	Number (%) Abnormal in 1997	Adj. Relative Risk (95% C.I.) ^{ab}	p-Value ^b
Comparison	1,017	23 (2.3)		
Background RH	353	8 (2.3)	0.87 (0.38,1.97)	0.731
Low RH	224	7 (3.1)	1.30 (0.55,3.09)	0.555
High RH	238	7 (2.9)	1.79 (0.75,4.30)	0.193
Low plus High RH	462	14 (3.0)	1.53 (0.77,3.03)	0.221

^a Relative risk and confidence interval relative to Comparisons.

Note: RH = Ranch Hand.

Comparison: $1987 \text{ Dioxin} \le 10 \text{ ppt.}$

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin >10 ppt, 10 ppt < Initial Dioxin ≤ 94 ppt.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

Summary statistics for 1992 are provided for reference purposes for participants who attended the 1985 and 1997 examinations. Statistical analyses are based only on participants who had normal populates in 1985 (see Chapter 7, Statistical Methods).

14.2.3.1.5 Dorsalis Pedis Pulses

The longitudinal analyses in Models 1 through 3 did not reveal any significant associations between dioxin and the change in dorsalis pedis pulses (Table 14-35(a–c): p>0.33 for each analysis).

Table 14-35. Longitudinal Analysis of Dorsalis Pedis Pulses

Occupational	Number (%) Abnormal/(n) Examination					
Category	Group	1985	1992	1997		
All	Ranch Hand	94 (11.4) (821)	60 (7.5) (798)	67 (8.2) (821)		
	Comparison	111 (10.6) (1,044)	70 (6.9) (1,017)	85 (8.1) (1,044)		
Officer	Ranch Hand	41 (12.9) (318)	23 (7.4) (312)	27 (8.5) (318)		
	Comparison	43 (10.5) (409)	28 (7.0) (402)	30 (7.3) (409)		
Enlisted Flyer	Ranch Hand	16 (11.0) (145)	9 (6.3) (143)	18 (12.4) (145)		
	Comparison	23 (14.6) (158)	16 (10.3) (156)	13 (8.2) (158)		
Enlisted Groundcrew	Ranch Hand	37 (10.3) (358)	28 (8.2) (343)	22 (6.1) (358)		
	Comparison	45 (9.4) (477)	26 (5.7) (459)	42 (8.8) (477)		

^b Adjusted for percent body fat at the time of the blood measurement of dioxin and age in 1997.

Table 14-35. Longitudinal Analysis of Dorsalis Pedis Pulses (Continued)

	_	Normal in 1985				
Occupational Category	Group	n in 1997	Number (%) Abnormal in 1997	Adj. Relative Risk (95% C.I.) ^a	p-Value ^a	
All	Ranch Hand Comparison	727 933	50 (6.9) 66 (7.1)	0.97 (0.66,1.43)	0.894	
Officer	Ranch Hand Comparison	277 366	22 (7.9) 27 (7.4)	1.07 (0.59,1.93)	0.821	
Enlisted Flyer	Ranch Hand Comparison	129 135	12 (9.3) 9 (6.7)	1.42 (0.58,3.52)	0.444	
Enlisted Groundcrew	Ranch Hand Comparison	321 432	16 (5.0) 30 (6.9)	0.73 (0.39,1.38)	0.335	

^a Relative risk, confidence interval, and p-values are in reference to a contrast of 1985 and 1997 results; results adjusted for age in 1997.

Note: Summary statistics for 1992 are provided for reference purposes for participants who attended the 1985 and 1997 examinations. Statistical analyses are based only on participants who had normal dorsalis pedis pulses in 1985 (see Chapter 7, Statistical Methods).

(b) MODEL 2: RANCH HANDS — INITIAL DIOXIN	1
---	---

		Number (%) Abnormal/(n) Examination	
Initial Dioxin	1985	1992	1997
Low	14 (9.4)	8 (5.6)	12 (8.1)
	(149)	(144)	(149)
Medium	20 (12.7)	14 (9.0)	16 (10.1)
	(158)	(155)	(158)
High	12 (7.8)	9 (6.0)	10 (6.5)
-	(154)	(149)	(154)

Initial	Dioxin Category Su	ımmary Statistics	Analysis Results for Log ₂ (Initial Dioxin) ^a		
Normal in 1985					
Initial Dioxin	n in 1997	Number (%) Abnormal in 1997	Adj. Relative Risk (95% C.I.) ^b	p-Value	
Low	135	10 (7.4)	1.01 (0.72,1.41)	0.946	
Medium	138	11 (8.0)			
High	142	7 (4.9)			

^a Adjusted for percent body fat at the time of the blood measurement of dioxin and age in 1997.

Note: Low = 27-63 ppt; Medium = >63-152 ppt; High = >152 ppt.

Summary statistics for 1992 are provided for reference purposes for participants who attended the 1985 and 1997 examinations. Statistical analyses are based only on participants who had normal dorsalis pedis pulses in 1985 (see Chapter 7, Statistical Methods).

b Relative risk for a twofold increase in initial dioxin.

Table 14-35. Longitudinal Analysis of Dorsalis Pedis Pulses (Continued)

(c) MODEL 3.	RANCH HANDS	AND COMP	ARISONS BY DIOXIN	CATECORY
TO MICHIEL 5.	NAMEDIAM	A Y 7 7 7 7 7 7 7 7 7	ANISONS DI 1710/AUS	CALINTIAL

	Number (%) Abnormal/(n) Examination			
Dioxin Category	1985	1992	1997	
Comparison	108 (10.6)	70 (7.1)	85 (8.4)	
_	(1,016)	(991)	(1,016)	
Background RH	48 (13.5)	29 (8.4)	29 (8.2)	
	(355)	(345)	(355)	
Low RH	21 (9.4)	12 (5.5)	22 (9.8)	
	(224)	(217)	(224)	
High RH	25 (10.5)	19 (8.2)	16 (6.8)	
•	(237)	(231)	(237)	
Low plus High RH	46 (10.0)	31 (6.9)	38 (8.2)	
1 0	(461)	(448)	(461)	

	Norm	nal in 1985		
Dioxin Category	n in 1997	Number (%) Abnormal in 1997	Adj. Relative Risk (95% C.I.) ^{ab}	p-Value ^b
Comparison	908	66 (7.3)		
Background RH	307	22 (7.2)	0.89 (0.53,1.48)	0.650
Low RH	203	17 (8.4)	1.08 (0.61,1.89)	0.798
High RH	212	11 (5.2)	0.91 (0.47,1.78)	0.789
Low plus High RH	415	28 (6.7)	0.99 (0.62,1.59)	0.964

^a Relative risk and confidence interval relative to Comparisons.

Comparison: $1987 \text{ Dioxin} \le 10 \text{ ppt.}$

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin >10 ppt, 10 ppt < Initial Dioxin ≤ 94 ppt.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

Summary statistics for 1992 are provided for reference purposes for participants who attended the 1985 and 1997 examinations. Statistical analyses are based only on participants who had normal dorsalis pedis pulses in 1985 (see Chapter 7, Statistical Methods).

14.2.3.1.6 Posterior Tibial Pulses

Model 1 and 2 analyses did not show any significant associations between dioxin and the change in posterior tibial pulses between 1985 and 1997 (Table 14-36(a,b): p>0.12 for each analysis).

Model 3 analysis of the change in posterior tibial pulses from normal in 1985 to abnormal in 1997 revealed one significant and one marginally significant contrast: Ranch Hands in the high dioxin category versus Comparisons (Table 14-36(c): Adj. RR=1.70, p=0.090) and Ranch Hands in the low plus high dioxin category versus Comparisons (Table 14-36(c): Adj. RR=1.60, p=0.047). Of the Comparisons, 5.1 percent had normal posterior tibial pulses in 1985 and abnormal posterior tibial pulses

^b Adjusted for percent body fat at the time of the blood measurement of dioxin and age in 1997.

in 1997. Of the Ranch Hands, 6.3 percent in the high dioxin category and 7.2 percent in the low plus high dioxin category had normal posterior tibial pulses in 1985 and abnormal posterior tibial pulses in 1997.

Table 14-36. Longitudinal Analysis of Posterior Tibial Pulses

Occupational		Number (%) Abnormal/(n) Examination				
Category	Group	1985	1992	1997		
A <i>ll</i>	Ranch Hand	2 (0.2) (822)	20 (2.5) (801)	56 (6.8) (822)		
	Comparison	6 (0.6) (1,044)	22 (2.2) (1,017)	58 (5.6) (1,044)		
Officer	Ranch Hand	1 (0.3) (318)	9 (2.9) (313)	21 (6.6) (318)		
	Comparison	2 (0.5) (411)	10 (2.5) (404)	23 (5.6) (411)		
nlisted Flyer	Ranch Hand	1 (0.7) (145)	5 (3.5) (143)	14 (9.7) (145)		
	Comparison	1 (0.6) (156)	4 (2.6) (154)	10 (6.4) (156)		
nlisted Groundcrew	Ranch Hand	0 (0.0) (359)	6 (1.7) (345)	21 (5.8) (359)		
	Comparison	3 (0.6)	8 (1.7)	25 (5.2)		

		Nor	mal in 1985		
Occupational Category	Group	n in 1997	Number (%) Abnormal in 1997	Adj. Relative Risk (95% C.I.) ^a	p-Value ^a
All	Ranch Hand Comparison	820 1,038	56 (6.8) 53 (5.1)	1.36 (0.92,2.01)	0.129
Officer	Ranch Hand Comparison	317 409	21 (6.6) 21 (5.1)	1.29 (0.69,2.43)	0.423
Enlisted Flyer	Ranch Hand Comparison	144 155	14 (9.7) 9 (5.8)	1.70 (0.70,4.09)	0.239
Enlisted Groundcrew	Ranch Hand Comparison	359 474	21 (5.8) 23 (4.9)	1.26 (0.68,2.35)	0.458

(477)

(459)

(477)

Note: Summary statistics for 1992 are provided for reference purposes for participants who attended the 1985 and 1997 examinations. Statistical analyses are based only on participants who had normal posterior tibial pulses in 1985 (see Chapter 7, Statistical Methods).

^a Relative risk, confidence interval, and p-values are in reference to a contrast of 1985 and 1997 results; results adjusted for age in 1997.

Table 14-36. Longitudinal Analysis of Posterior Tibial Pulses (Continued)

(b) MODEL 2: RAN	CH HANDS — INITIAL 1	DIOXIN	
		Number (%) Abnormal/(n) Examination	
Initial Dioxin	1985	1992	1997
Low	1 (0.7)	5 (3.5)	9 (6.0)
	(149)	(144)	(149)
Medium	0 (0.0)	5 (3.2)	15 (9.5)
	(158)	(155)	(158)
High	1 (0.6)	2(1.3)	9 (5.8)
-	(155)	(151)	(155)

Initial	Dioxin Category St	ımmary Statistics	Analysis Results for Log	(Initial Dioxin) ^a
	No	rmal in 1985		
Initial Dioxin	n in 1997	Number (%) Abnormal in 1997	Adj. Relative Risk (95% C.I.) ^b	p-Value
Low	148	9 (6.1)	1.12 (0.85,1.49)	0.418
Medium	158	15 (9.5)		
High	154	9 (5.8)		

^a Adjusted for percent body fat at the time of the blood measurement of dioxin and age in 1997.

Note: Low = 27-63 ppt; Medium = >63-152 ppt; High = >152 ppt.

Summary statistics for 1992 are provided for reference purposes for participants who attended the 1985 and 1997 examinations. Statistical analyses are based only on participants who had normal posterior tibial pulses in 1985 (see Chapter 7, Statistical Methods).

(c) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY

		Number (%) Abnormal/(n) Examination	
Dioxin Category	1985	1992	1997
Comparison	6 (0.6)	22 (2.2)	57 (5.6)
•	(1,016)	(991)	(1,016)
Background RH	0 (0.0)	7 (2.0)	22 (6.2)
C	(355)	(346)	(355)
Low RH	1 (0.4)	6 (2.8)	18 (8.0)
	(224)	(217)	(224)
High RH	1 (0.4)	6 (2.6)	15 (6.3)
C	(238)	(233)	(238)
Low plus High RH	2 (0.4)	12 (2.7)	33 (7.1)
	(462)	(450)	(462)

^b Relative risk for a twofold increase in initial dioxin.

Table 14-36. Longitudinal Analysis of Posterior Tibial Pulses (Continued)

	Norm	nal in 1985		
Dioxin Category	n in 1997	Number (%) Abnormal in 1997	Adj. Relative Risk (95% C.I.) ^{ab}	p-Value ^b
Comparison	1,010	52 (5.1)		
Background RH	355	22 (6.2)	1.05 (0.62,1.77)	0.856
Low RH	223	18 (8.1)	1.50 (0.85, 2.65)	0.160
High RH	237	15 (6.3)	1.70 (0.92,3.12)	0.090
Low plus High RH	460	33 (7.2)	1.60 (1.01, 2.54)	0.047

^a Relative risk and confidence interval relative to Comparisons.

Comparison: $1987 \text{ Dioxin} \le 10 \text{ ppt.}$

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin > 10 ppt, $10 \text{ ppt} < \text{Initial Dioxin} \le 94 \text{ ppt}$.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

Summary statistics for 1992 are provided for reference purposes for participants who attended the 1985 and 1997 examinations. Statistical analyses are based only on participants who had normal posterior tibial pulses in 1985 (see Chapter 7, Statistical Methods).

14.2.3.1.7 Leg Pulses

The longitudinal analyses in Models 1 through 3 did not reveal a significant association between dioxin and the change from normal leg pulses in 1985 to abnormal leg pulses in 1997 (Table 14-37(a–c): p>0.15 for each analysis).

Table 14-37. Longitudinal Analysis of Leg Pulses

Occupational	_	ľ	Number (%) Abnormal/(n) Examination		
Category	Group	1985	1992	1997	
All	Ranch Hand	97 (11.8) (821)	66 (8.3) (798)	91 (11.1) (821)	
	Comparison	114 (10.9) (1,042)	77 (7.6) (1,015)	109 (10.5) (1,042)	
Officer	Ranch Hand	43 (13.5) (318)	24 (7.7) (312)	35 (11.0) (318)	
	Comparison	44 (10.8) (409)	29 (7.2) (402)	38 (9.3) (409)	
Enlisted Flyer	Ranch Hand	17 (11.7) (145)	11 (7.7) (143)	25 (17.2) (145)	
	Comparison	22 (14.1) (156)	16 (10.4) (154)	17 (10.9) (156)	
Enlisted Groundcrew	Ranch Hand	37 (10.3) (358)	31 (9.0) (343)	31 (8.7) (358)	
	Comparison	48 (10.1) (477)	32 (7.0) (459)	54 (11.3) (477)	

^b Adjusted for percent body fat at the time of the blood measurement of dioxin and age in 1997.

Table 14-37. Longitudinal Analysis of Leg Pulses (Continued)

		Nor	mal in 1985		
Occupational Category	Group	n in 1997	Number (%) Abnormal in 1997	Adj. Relative Risk (95% C.I.) ^a	p-Value ^a
All	Ranch Hand Comparison	724 928	73 (10.1) 85 (9.2)	1.12 (0.80,1.57)	0.502
Officer	Ranch Hand Comparison	275 365	29 (10.5) 34 (9.3)	1.13 (0.67,1.93)	0.645
Enlisted Flyer	Ranch Hand Comparison	128 134	19 (14.8) 12 (9.0)	1.76 (0.81,3.83)	0.153
Enlisted Groundcrew	Ranch Hand Comparison	321 429	25 (7.8) 39 (9.1)	0.89 (0.52,1.52)	0.676

^a Relative risk, confidence interval, and p-values are in reference to a contrast of 1985 and 1997 results; results adjusted for age in 1997.

Note: Summary statistics for 1992 are provided for reference purposes for participants who attended the 1985 and 1997 examinations. Statistical analyses are based only on participants who had normal leg pulses in 1985 (see Chapter 7, Statistical Methods).

(b) MODEL 2: RANCH HANDS — INITIAL DIOXIN

	Number (%) Abnormal/(n) Examination		
Initial Dioxin	1985	1992	1997
W	15 (10.1)	9 (6.3)	15 (10.1)
	(149)	(144)	(149)
edium	20 (12.7)	17 (11.0)	22 (13.9)
	(158)	(155)	(158)
gh	13 (8.4)	9 (6.0)	14 (9.1)
•	(154)	(149)	(154)

Initial Dioxin Category Summary Statistics			Analysis Results for Log	₂ (Initial Dioxin) ^a
	No	rmal in 1985		
Initial Dioxin	n in 1997	Number (%) Abnormal in 1997	Adj. Relative Risk (95% C.I.) ^b	p-Value
Low	134	13 (9.7)	1.14 (0.87,1.49)	0.344
Medium	138	17 (12.3)		
High	141	11 (7.8)		

^a Adjusted for percent body fat at the time of the blood measurement of dioxin and age in 1997.

Note: Low = 27-63 ppt; Medium = >63-152 ppt; High = >152 ppt.

Summary statistics for 1992 are provided for reference purposes for participants who attended the 1985 and 1997 examinations. Statistical analyses are based only on participants who had normal leg pulses in 1985 (see Chapter 7, Statistical Methods).

^b Relative risk for a twofold increase in initial dioxin.

Table 14-37. Longitudinal Analysis of Leg Pulses (Continued)

() MODEL 4 DANGER HAND AND COMPAND TO SERVED TO SERVE OF THE CORE
(c) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY

		Number (%) Abnormal/(n) Examination	
Dioxin Category	1985	1992	1997
Comparison	111 (10.9)	77 (7.8)	108 (10.7)
_	(1,014)	(989)	(1,014)
Background RH	49 (13.8)	30 (8.7)	39 (11.0)
	(355)	(345)	(355)
Low RH	22 (9.8)	13 (6.0)	29 (12.9)
	(224)	(217)	(224)
High RH	26 (11.0)	22 (9.5)	22 (9.3)
C	(237)	(231)	(237)
Low plus High RH	48 (10.4)	35 (7.8)	51 (11.1)
	(461)	(448)	(461)

_	Normal in 1985 Number (%) n in 1997 Abnormal in 1997			
Dioxin Category			Adj. Relative Risk (95% C.I.) ^{ab}	p-Value ^b
Comparison	903	84 (9.3)		
Background RH	306	31 (10.1)	0.98 (0.63,1.52)	0.924
Low RH	202	24 (11.9)	1.21 (0.74,1.97)	0.455
High RH	211	17 (8.1)	1.17 (0.67,2.04)	0.589
Low plus High RH	413	41 (9.9)	1.19 (0.79,1.78)	0.411

^a Relative risk and confidence interval relative to Comparisons.

Comparison: $1987 \text{ Dioxin} \le 10 \text{ ppt.}$

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin >10 ppt, 10 ppt < Initial Dioxin ≤ 94 ppt.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

Summary statistics for 1992 are provided for reference purposes for participants who attended the 1985 and 1997 examinations. Statistical analyses are based only on participants who had normal leg pulses in 1985 (see Chapter 7, Statistical Methods).

14.2.3.1.8 Peripheral Pulses

The change from normal peripheral pulses in 1985 to abnormal peripheral pulses in 1997 was not significantly associated with dioxin in Models 1 through 3 (Table 14-38(a–c): p>0.15 for each analysis).

^b Adjusted for percent body fat at the time of the blood measurement of dioxin and age in 1997.

Table 14-38. Longitudinal Analysis of Peripheral Pulses

(a) MODEL 1: RANCH HANDS VS. COMPARISONS

Occupational]	Number (%) Abnormal/(r Examination	1)
Category	Group	1985	1992	1997
All	Ranch Hand	97 (11.8) (821)	66 (8.3) (798)	94 (11.4) (821)
	Comparison	116 (11.1) (1,041)	81 (8.0) (1,014)	112 (10.8) (1,041)
Officer	Ranch Hand	43 (13.5) (318)	24 (7.7) (312)	36 (11.3) (318)
	Comparison	44 (10.8) (409)	30 (7.5) (402)	40 (9.8) (409)
Enlisted Flyer	Ranch Hand	17 (11.7) (145)	11 (7.7) (143)	25 (17.2) (145)
	Comparison	22 (14.1) (156)	16 (10.4) (154)	17 (10.9) (156)
Enlisted Groundcrew	Ranch Hand	37 (10.3) (358)	31 (9.0) (343)	33 (9.2) (358)
	Comparison	50 (10.5) (476)	35 (7.6) (458)	55 (11.6) (476)

	_	Normal in 1985			
Occupational Category	Group	n in 1997	Number (%) Abnormal in 1997	Adj. Relative Risk (95% C.I.) ^a	p-Value ^a
All	Ranch Hand Comparison	724 925	76 (10.5) 87 (9.4)	1.14 (0.82,1.59)	0.433
Officer	Ranch Hand Comparison	275 365	30 (10.9) 36 (9.9)	1.10 (0.66,1.86)	0.710
Enlisted Flyer	Ranch Hand Comparison	128 134	19 (14.8) 12 (9.0)	1.76 (0.81,3.83)	0.154
Enlisted Groundcrew	Ranch Hand Comparison	321 426	27 (8.4) 39 (9.2)	0.97 (0.57,1.64)	0.901

^a Relative risk, confidence interval, and p-values are in reference to a contrast of 1985 and 1997 results; results adjusted for age in 1997.

Note: Summary statistics for 1992 are provided for reference purposes for participants who attended the 1985 and 1997 examinations. Statistical analyses are based only on participants who had normal peripheral pulses in 1985 (see Chapter 7, Statistical Methods).

Table 14-38. Longitudinal Analysis of Peripheral Pulses (Continued)

(h) MODEL	2:	RANCH	HANDS —	INITIAL	DIOXIN

		Number (%) Abnormal/(n) Examination	
Initial Dioxin	1985	1992	1997
Low	15 (10.1)	9 (6.3)	16 (10.7)
	(149)	(144)	(149)
Medium	20 (12.7)	17 (11.0)	22 (13.9)
	(158)	(155)	(158)
High	13 (8.4)	9 (6.0)	15 (9.7)
	(154)	(149)	(154)

Initial Dioxin Category Summary Statistics			Analysis Results for Log	₂ (Initial Dioxin) ^a
Normal in 1985				
Initial Dioxin	n in 1997	Number (%) Abnormal in 1997	Adj. Relative Risk (95% C.I.) ^b	p-Value
Low	134	14 (10.4)	1.11 (0.85,1.45)	0.434
Medium	138	17 (12.3)		
High	141	12 (8.5)		

^a Adjusted for percent body fat at the time of the blood measurement of dioxin and age in 1997. ^b Relative risk for a twofold increase in initial dioxin.

Note: Low = 27-63 ppt; Medium = >63-152 ppt; High = >152 ppt.

Summary statistics for 1992 are provided for reference purposes for participants who attended the 1985 and 1997 examinations. Statistical analyses are based only on participants who had normal peripheral pulses in 1985 (see Chapter 7, Statistical Methods).

(c) MODEL 3: RANCH HANDS AND COMPARISONS BY DIOXIN CATEGORY

	Number (%) Abnormal/(n) Examination				
Dioxin Category	1985	1992	1997		
Comparison	113 (11.2)	81 (8.2)	111 (11.0)		
-	(1,013)	(988)	(1,013)		
Background RH	49 (13.8)	30 (8.7)	40 (11.3)		
-	(355)	(345)	(355)		
Low RH	22 (9.8)	13 (6.0)	30 (13.4)		
	(224)	(217)	(224)		
High RH	26 (11.0)	22 (9.5)	23 (9.7)		
	(237)	(231)	(237)		
Low plus High RH	48 (10.4)	35 (7.8)	53 (11.5)		
	(461)	(448)	(461)		

Table 14-38. Longitudinal Analysis of Peripheral Pulses (Continued)

	Normal in 1985			
Dioxin Category	n in 1997	Number (%) Abnormal in 1997	Adj. Relative Risk (95% C.I.) ^{ab}	p-Value ^b
Comparison	900	86 (9.6)		
Background RH	306	32 (10.5)	0.98 (0.63,1.52)	0.934
Low RH	202	25 (12.4)	1.23 (0.76,1.99)	0.408
High RH	211	18 (8.5)	1.22 (0.70,2.11)	0.482
Low plus High RH	413	43 (10.4)	1.22 (0.82,1.82)	0.325

^a Relative risk and confidence interval relative to Comparisons.

Comparison: $1987 \text{ Dioxin} \le 10 \text{ ppt.}$

Background (Ranch Hand): 1987 Dioxin ≤ 10 ppt.

Low (Ranch Hand): 1987 Dioxin >10 ppt, 10 ppt < Initial Dioxin ≤ 94 ppt.

High (Ranch Hand): 1987 Dioxin > 10 ppt, Initial Dioxin > 94 ppt.

Summary statistics for 1992 are provided for reference purposes for participants who attended the 1985 and 1997 examinations. Statistical analyses are based only on participants who had normal peripheral pulses in 1985 (see Chapter 7, Statistical Methods).

14.3 DISCUSSION

Cardiovascular diseases are among the most common encountered by the primary care physician. In practice, the diagnosis of cardiovascular disease is based primarily on the noninvasive data analyzed in the current chapter. Specifically, the history, physical examination, chest x ray, and resting ECG remain highly reliable indices that can alert the clinician to the presence of underlying cardiovascular disease and indicate the need for additional, more specific, noninvasive or invasive studies. Although arbitrary, dividing data collection into central and peripheral cardiovascular functions is convenient and forms a reasonable basis for comparison of the cohorts under study.

The limitations of the history in cardiovascular diagnosis deserve emphasis. In peripheral vascular disease, for example, signs and symptoms will vary depending on the degree of development of collateral circulatory channels. While hemodynamically significant arterial disease of the lower extremities is usually associated with claudication, severe carotid occlusive disease can be present in the absence of symptoms of transient cerebral ischemia. Further, conclusive evidence shows that advanced coronary artery disease can occur in the absence of angina and be present as "silent" myocardial ischemia. Lastly, it is well recognized that the cardiovascular history, as related by patients, is often subject to error. The generic term "heart attack," for example, can be used to describe any type of cardiac event from an isolated episode of unstable angina or arrhythmia to a myocardial infarction. These imperfections highlight the importance of the medical record verification conducted in this study.

In the cardiovascular assessment particularly, the physical examination can provide valuable clues to the presence of asymptomatic but significant underlying disease. Steps were taken to simplify data collection and reduce differences among the examining physicians. All blood pressure readings, for example, were taken by automated sphygmomanometric instruments. Auscultory endpoints—murmurs and bruits—were recorded as present or absent by anatomic location, thus eliminating speculation as to specific

^b Adjusted for percent body fat at the time of the blood measurement of dioxin and age in 1997.

valvular or vessel origin and hemodynamic significance. As markers of occult arterial occlusive disease, vascular bruits are relatively easy to detect and were carefully sought over the carotid, abdominal, and femoral vessels.

The data relevant to this chapter included the resting ECG, the standard two-view chest x ray (discussed in Chapter 18, Pulmonary Assessment) and Doppler arterial vascular studies. The test used can confirm diagnoses that can be made based on data available in the current assessment. For example, when correlated with the history and physical examination, the chest x ray and ECG enable the clinician to draw highly accurate conclusions regarding the presence and hemodynamic significance of valvular heart disease of any etiology. As defined by the chest x ray, the pulmonary vascularity can provide reliable clues to the presence of global left ventricular dysfunction with pulmonary venous congestion and of pulmonary hypertension of any cause.

In the analyses of verified historical variables, hypertension, myocardial infarction, transient ischemic attack, and stroke were similar in Ranch Hands and Comparisons. In the 1997 examinations, in contrast to 1992, Ranch Hands were more likely to have a history of heart disease (66.1% vs. 60.8%) across all occupational strata, particularly in the enlisted flyer category. In none of the physical examination or electrocardiographic variables were any significant group differences defined. The prevalence of funduscopic abnormalities, peripheral pulse deficits, and intermittent claudication, all more common in Ranch Hands than Comparisons in the 1992 examination, is now essentially the same in the two cohorts.

Serum dioxin analyses yielded several significant results. In the unadjusted analysis, a significant positive dose-response effect was noted in Ranch Hands in the association of hypertension with 1987 serum dioxin levels (34.0%, 38.0%, and 49.1% in the low, medium, and high categories, respectively), an association that remained significant after adjustment for covariates. Similarly, although the association was less significant, a positive dose-response effect was noted between the electrocardiographic evidence of a myocardial infarction and both initial and 1987 serum dioxin levels. Ranch Hands in the highest dioxin category were more likely than Comparisons to have tachycardia, as determined by the electrocardiograph. In contrast, although Ranch Hands were more likely than Comparisons to have a history of heart disease, a significant inverse dose-response effect was noted in relation to both extrapolated initial and 1987 serum dioxin levels. These results are consistent with those from both the 1987 and 1992 examinations.

With few exceptions, dependent variable-covariate analyses confirmed well-established associations. By a medical records review and by abnormalities detected on physical examinations, cardiovascular disease was associated significantly with the classic risk factors of age, cigarette use, and, particularly, diabetes. Obesity proved to be a significant risk factor for the development of heart disease and for numerous electrocardiographic abnormalities but not to the occurrence of myocardial infarction historically or by ECG. Alcohol consumption was associated strongly with the development of hypertension but did not have the protective effect on the occurrence of myocardial infarction that was noted in the 1992 examination. The increased prevalence of pulse deficits in association with alcohol consumption may have been mediated by concomitant cigarette use. Finally, consistent with the results of the 1987 and 1992 examinations, type A personality traits were not found to be associated with an increased risk for the development of cardiovascular disease.

In the longitudinal analysis, a comparable increase in the prevalence of peripheral pulse deficits was noted in both the Ranch Hand and Comparison cohorts between the 1992 and 1997 examinations. Although none of the group differences was statistically significant, Ranch Hands continued to have a slightly greater prevalence of pulse deficits than Comparisons at all sites examined. Two of the six analyses, the posterior tibial and femoral pulses, yielded evidence for a significant or marginally significant association

of pulse deficits with categorized dioxin. Consistent with all previous examinations, Comparisons were found to be at slightly greater risk than Ranch Hands for the development of systolic hypertension by discrete analysis, but group differences remain nonsignificant.

In contrast to prior examinations, the current study has documented that Ranch Hands are more likely than Comparisons to have historical evidence for heart disease (excluding essential hypertension) but are no longer at greater risk for the occurrence of pulse deficits. By all other indices, the prevalence of cardiovascular disease appears similar in both cohorts. For the first time, there is evidence that dioxin exposure may be a risk factor for the development of hypertension and myocardial infarction. As of 1997, the verified history of essential hypertension was associated with 1987 dioxin, and the evidence of prior myocardial infarction from the ECG was associated with initial dioxin.

14.4 SUMMARY

The cardiovascular assessment was based on a medical records review and verification, physical examination and ECG determinations, and an ICVI index based on participant responses to three questions regarding leg pain. Variables constructed from the medical records review included essential hypertension, heart disease (excluding essential hypertension), myocardial infarction, and stroke or transient ischemic attack. The physical examination findings, the ECG determinations, and the ICVI index investigated the central cardiac function and peripheral vascular function. Each health endpoint was examined for an association with exposure group (Model 1), initial dioxin (Model 2), categorized dioxin (Model 3), and 1987 dioxin levels (Model 4). Significant results from the adjusted analyses are presented below.

14.4.1 Model 1: Group Analysis

The adjusted group analysis revealed that Ranch Hands had a significantly higher percentage of participants with a history of heart disease (excluding essential hypertension) than did Comparisons when all occupational strata were combined. Stratifying by occupation revealed a significantly higher percentage of Ranch Hand enlisted flyers with a history of heart disease than Comparison enlisted flyers. Ranch Hand enlisted groundcrew had a significantly lower percentage of abnormal funduscopic examination results than Comparison enlisted groundcrew. Ranch Hand enlisted groundcrew also had a marginally significantly lower percentage of abnormal overall ECG findings than Comparison enlisted groundcrew. The results of all unadjusted and adjusted Model 1 analyses are summarized in Table 14-39.

Table 14-39. Summary of Group Analysis (Model 1) for Cardiovascular Variables (Ranch Hands vs. Comparisons)

	UNADJUSTED				
Variable	All	Officer	Enlisted Flyer	Enlisted Groundcrew	
Medical Records					
Essential Hypertension (D)	ns	ns	NS	ns	
Heart Disease (Excluding Essential Hypertension) (D)	+0.013	NS	+0.003	NS	
Myocardial Infarction (D)	NS	ns	NS	NS	
Stroke or Transient Ischemic Attack (D)	NS	NS	ns	NS	
Physical Examination					
Systolic Blood Pressure (C)	ns	ns	ns	ns	
Systolic Blood Pressure (D)	ns	NS	NS	ns	

Table 14-39. Summary of Group Analysis (Model 1) for Cardiovascular Variables (Ranch Hands vs. Comparisons) (Continued)

	UNADJUSTED			
Variable	All	Officer	Enlisted Flyer	Enlisted Groundcrew
Diastolic Blood Pressure (C)	ns	ns	NS	ns
Diastolic Blood Pressure (D)	NS	NS	NS	ns
Heart Sounds (D)	ns	ns	ns	ns
Overall Electrocardiograph (ECG) (D)	NS	NS	NS	ns*
ECG: Right Bundle Branch Block (D)	ns	ns	NS	ns
ECG: Left Bundle Branch Block (D)	ns	ns	NS	ns
ECG: Non-Specific ST- and T-Wave Changes (D)	NS	NS	NS	ns
ECG: Bradycardia (D)	ns	ns	NS	ns
ECG: Tachycardia (D)	NS	NS	NS	NS
ECG: Arrhythmia (D)	NS	NS	NS	ns
ECG: Evidence of Prior Myocardial Infarction (D)	ns	ns	ns	ns
ECG: Other Diagnoses (D)	NS	NS		NS
Funduscopic Examination (D)	ns	NS	NS	-0.033
Carotid Bruits (D)	NS	ns	NS	ns
Radial Pulses (D)	NS	NS		NS
Femoral Pulses (D)	NS*	NS	NS	NS
Popliteal Pulses (D)	NS	ns	NS	NS
Dorsalis Pedis Pulses (D)	NS	NS	NS	ns
Posterior Tibial Pulses (D)	NS	NS	NS	NS
Leg Pulses (D)	NS	NS	NS	ns
Peripheral Pulses (D)	NS	NS	NS	ns
Self-reported Questionnaire				
Intermittent Claudication and Vascular Insufficiency Index (ICVI) (D)	NS	NS	ns	NS

NS* or ns*: Marginally significant (0.05<p≤0.10).

C: Continuous analysis.

D: Discrete analysis.

+: Relative risk ≥ 1.00 .

-: Relative risk <1.00.

--: Analysis not performed because of the sparse number of participants with an abnormality.

P-value given if $p \le 0.05$.

A capital "NS" denotes a relative risk of 1.00 or greater for discrete analysis or differences of means nonnegative for continuous analysis. A lowercase "ns" denotes a relative risk less than 1.00 for discrete analysis or difference of means negative for continuous analysis.

Table 14-39. Summary of Group Analysis (Model 1) for Cardiovascular Variables (Ranch Hands vs. Comparisons) (Continued)

	ADJUSTED			
Variable	All	Officer	Enlisted Flyer	Enlisted Groundcrew
Medical Records				
Essential Hypertension (D)	ns	ns	NS	ns
Heart Disease (Excluding Essential Hypertension) (D)	+0.018	NS	+0.004	NS
Myocardial Infarction (D)	NS	ns	NS	ns
Stroke or Transient Ischemic Attack (D)	NS	NS		NS
Physical Examination				
Systolic Blood Pressure (C)	ns	ns	NS	ns
Systolic Blood Pressure (D)	ns	ns	NS	ns
Diastolic Blood Pressure (C)	NS	ns	NS	NS
Diastolic Blood Pressure (D)	NS	NS	NS	ns
Heart Sounds (D)	ns	ns	ns	ns
Overall Electrocardiograph (ECG) (D)	ns	NS	NS	ns*
ECG: Right Bundle Branch Block (D)	ns	ns	NS	ns
ECG: Left Bundle Branch Block (D)	ns	ns		ns
ECG: Non-Specific ST- and T-Wave Changes (D)	NS	NS	NS	ns
ECG: Bradycardia (D)	ns	ns	NS	ns
ECG: Tachycardia (D)	NS			NS
ECG: Arrhythmia (D)	NS	NS	NS	ns
ECG: Evidence of Prior Myocardial Infarction (D)	ns	ns	NS	ns
ECG: Other Diagnoses (D)	NS			NS
Funduscopic Examination (D)	ns	NS	NS	-0.047
Carotid Bruits (D)	ns	ns	NS	ns
Radial Pulses (D)	NS	NS		NS
Femoral Pulses (D)	NS	NS	NS	NS
Popliteal Pulses (D)	NS	ns	ns	NS
Dorsalis Pedis Pulses (D)	ns	NS	NS	ns
Posterior Tibial Pulses (D)	NS	NS	NS	NS
Leg Pulses (D)	NS	NS	NS	ns
Peripheral Pulses (D)	NS	NS	NS	ns
Self-reported Questionnaire				
Intermittent Claudication and Vascular Insufficiency	ns	NS	ns	NS
Index (ICVI) (D)				

ns*: Marginally significant (0.05<p≤0.10).

C: Continuous analysis.

- D: Discrete analysis.
- +: Relative risk ≥ 1.00 .
- -: Relative risk <1.00.
- --: Analysis not performed because of the sparse number of participants with an abnormality.

P-value given if $p \le 0.05$.

A capital "NS" denotes a relative risk of 1.00 or greater for discrete analysis or differences of means nonnegative for continuous analysis. A lowercase "ns" denotes a relative risk less than 1.00 for discrete analysis or difference of means negative for continuous analysis.

14.4.2 Model 2: Initial Dioxin Analysis

Model 2 analyses revealed a significant positive association between initial dioxin and evidence of prior myocardial infarction from the ECG. The results of all unadjusted and adjusted Model 2 analyses are summarized in Table 14-40.

Table 14-40. Summary of Initial Dioxin Analysis (Model 2) for Cardiovascular Variables (Ranch Hands Only)

Variable	Unadjusted	Adjusted
Medical Records		
Essential Hypertension (D)	NS	NS
Heart Disease (Excluding Essential Hypertension) (D)	-0.001	ns
Myocardial Infarction (D)	NS	NS
Stroke or Transient Ischemic Attack (D)	NS	NS
Physical Examination		
Systolic Blood Pressure (C)	ns	ns
Systolic Blood Pressure (D)	-0.031	ns
Diastolic Blood Pressure (C)	NS	NS
Diastolic Blood Pressure (D)	NS	NS
Heart Sounds (D)	NS	NS
Overall Electrocardiograph (ECG) (D)	ns	NS
ECG: Right Bundle Branch Block (D)	ns	NS
ECG: Left Bundle Branch Block (D)	ns	
ECG: Non-Specific ST- and T-Wave Changes (D)	ns	NS
ECG: Bradycardia (D)	ns	ns
ECG: Tachycardia (D)	NS	
ECG: Arrhythmia (D)	ns	NS
ECG: Evidence of Prior Myocardial Infarction (D)	NS	+0.012
ECG: Other Diagnoses (D)	NS	
Funduscopic Examination (D)	ns	NS
Carotid Bruits (D)	NS	NS
Radial Pulses (D)	ns	
Femoral Pulses (D)	ns	NS
Popliteal Pulses (D)	ns	ns
Dorsalis Pedis Pulses (D)	ns	NS
Posterior Tibial Pulses (D)	NS	NS
Leg Pulses (D)	ns	NS
Peripheral Pulses (D)	ns	NS
Self-reported Questionnaire		
Intermittent Claudication and Vascular Insufficiency	ns	NS
Index (ICVI) (D)		

Note: NS or ns: Not significant (p>0.10).

- C: Continuous analysis.
- D: Discrete analysis.
- +: Relative risk ≥ 1.00 .
- -: Relative risk <1.00.
- --: Analysis not performed because of the sparse number of Ranch Hands with an abnormality.

P-value given if $p \le 0.05$.

A capital "NS" denotes a relative risk of 1.00 or greater for discrete analysis or slope nonnegative for continuous analysis. A lowercase "ns" denotes a relative risk less than 1.00 for discrete analysis or slope negative for continuous analysis.

14.4.3 Model 3: Categorized Dioxin Analysis

The adjusted Model 3 analysis revealed a significantly higher occurrence of heart disease for Ranch Hands in the background dioxin category than for Comparisons. A significantly lower prevalence of abnormal heart sounds was found for Ranch Hands in the background dioxin category than for Comparisons. The percentage of Ranch Hands in the low dioxin category with a history of heart disease was marginally significantly greater than Comparisons. The prevalence of Ranch Hands in the low dioxin category with abnormal ECG findings was marginally significantly smaller than Comparisons. Ranch Hands in the high dioxin category had a significantly greater prevalence of tachycardia and other ECG diagnoses than Comparisons. The results of all unadjusted and adjusted Model 3 analyses are summarized in Table 14-41.

Table 14-41. Summary of Categorized Dioxin Analysis (Model 3) for Cardiovascular Variables (Ranch Hands vs. Comparisons)

	UNADJUSTED			
Variable	Background Ranch Hands vs. Comparisons	Low Ranch Hands vs. Comparisons	High Ranch Hands vs. Comparisons	Low plus High Ranch Hands vs. Comparisons
Medical Records				
Essential Hypertension (D)	ns	ns	NS	NS
Heart Disease (Excluding Essential	+0.005	+0.011	ns	NS
Hypertension) (D)				
Myocardial Infarction (D)	ns	ns	NS	NS
Stroke or Transient Ischemic Attack (D)	NS	ns	NS	ns
Physical Examination				
Systolic Blood Pressure (C)	ns	NS	ns*	ns
Systolic Blood Pressure (D)	NS	NS	ns	ns
Diastolic Blood Pressure (C)	ns	ns	NS*	NS
Diastolic Blood Pressure (D)	ns	NS	NS	NS
Heart Sounds (D)	-0.047	ns	ns	ns
Overall Electrocardiograph (ECG) (D)	NS	ns	ns	ns
ECG: Right Bundle Branch Block (D)	ns	ns	NS	ns
ECG: Left Bundle Branch Block (D)	NS	ns	ns	ns
ECG: Non-Specific ST- and T-Wave	ns	NS	NS	NS
Changes (D)				
ECG: Bradycardia (D)	ns	ns	-0.042	-0.020
ECG: Tachycardia (D)	NS	ns	+0.033	NS
ECG: Arrhythmia (D)	ns	NS	ns	NS
ECG: Evidence of Prior Myocardial	ns	NS	ns	ns
Infarction (D)				
ECG: Other Diagnoses (D)	NS	ns	+0.042	NS
Funduscopic Examination (D)	ns	NS	ns	NS
Carotid Bruits (D)	ns	ns	NS	NS
Radial Pulses (D)	NS	NS	NS	NS
Femoral Pulses (D)	NS	NS	NS	NS*
Popliteal Pulses (D)	ns	NS	NS	NS
Dorsalis Pedis Pulses (D)	ns	NS	ns	NS
Posterior Tibial Pulses (D)	NS	NS	NS	NS
Leg Pulses (D)	ns	NS	NS	NS
Peripheral Pulses (D)	ns	NS	NS	NS

Table 14-41. Summary of Categorized Dioxin Analysis (Model 3) for Cardiovascular Variables (Ranch Hands vs. Comparisons) (Continued)

	UNADJUSTED			
Variable	Background Ranch Hands vs. Comparisons	Low Ranch Hands vs. Comparisons	High Ranch Hands vs. Comparisons	Low plus High Ranch Hands vs. Comparisons
Self-reported Questionnaire Intermittent Claudication and Vascular Insufficiency Index (ICVI) (D)	ns	NS	NS	NS

NS* or ns*: Marginally significant (0.05<p≤0.10).

C: Continuous analysis.

D: Discrete analysis.

+: Relative risk ≥ 1.00 .

-: Relative risk <1.00.

P-value given if $p \le 0.05$.

A capital "NS" denotes a relative risk of 1.00 or greater for discrete analysis or differences of means nonnegative for continuous analysis. A lowercase "ns" denotes a relative risk less than 1.00 for discrete analysis or difference of means negative for continuous analysis.

	ADJUSTED			
Variable	Background Ranch Hands vs. Comparisons	Low Ranch Hands vs. Comparisons	High Ranch Hands vs. Comparisons	Low plus High Ranch Hands vs. Comparisons
Medical Records				
Essential Hypertension (D)	ns	ns	NS	NS
Heart Disease (Excluding Essential	+0.032	NS*	NS	NS
Hypertension) (D)				
Myocardial Infarction (D)	ns	ns	NS	NS
Stroke or Transient Ischemic Attack (D)	ns	ns	NS	NS
Physical Examination				
Systolic Blood Pressure (C)	NS	ns	ns	ns
Systolic Blood Pressure (D)	NS	NS	ns	ns
Diastolic Blood Pressure (C)	ns	ns	NS	NS
Diastolic Blood Pressure (D)	ns	ns	NS	NS
Heart Sounds (D)	-0.041	ns	NS	ns
Overall Electrocardiograph (ECG) (D)	NS	ns*	NS	ns
ECG: Right Bundle Branch Block (D)	NS	ns	NS	ns
ECG: Left Bundle Branch Block (D)	ns	ns		
ECG: Non-Specific ST- and T-Wave	ns	ns	NS	NS
Changes (D)				
ECG: Bradycardia (D)	ns	ns	ns	ns*
ECG: Tachycardia (D)	NS		+0.032	
ECG: Arrhythmia (D)	ns	NS	NS	NS
ECG: Evidence of Prior Myocardial	ns	ns	NS	ns
Infarction (D)				
ECG: Other Diagnoses (D)	NS		+0.050	
Funduscopic Examination (D)	NS	ns	ns	ns
Carotid Bruits (D)	NS	ns	NS	ns
Radial Pulses (D)	NS	NS	NS	NS
Femoral Pulses (D)	NS	NS	NS	NS

Table 14-41. Summary of Categorized Dioxin Analysis (Model 3) for Cardiovascular Variables (Ranch Hands vs. Comparisons) (Continued)

	ADJUSTED			
Variable	Background Ranch Hands vs. Comparisons	Low Ranch Hands vs. Comparisons	High Ranch Hands vs. Comparisons	Low plus High Ranch Hands vs. Comparisons
Popliteal Pulses (D)	ns	NS	NS	NS
Dorsalis Pedis Pulses (D)	ns	ns	ns	ns
Posterior Tibial Pulses (D)	NS	NS	NS	NS
Leg Pulses (D)	NS	NS	ns	ns
Peripheral Pulses (D)	NS	NS	ns	NS
Self-reported Questionnaire				
Intermittent Claudication and Vascular Insufficiency Index (ICVI) (D)	ns	ns	NS	NS

NS* or ns*: Marginally significant (0.05<p≤0.10).

C: Continuous analysis.

D: Discrete analysis.

+: Relative risk ≥ 1.00 .

-: Relative risk <1.00.

--: Analysis not performed because of the sparse number of participants with an abnormality.

P-value given if $p \le 0.05$.

A capital "NS" denotes a relative risk of 1.00 or greater for discrete analysis or differences of means nonnegative for continuous analysis. A lowercase "ns" denotes a relative risk less than 1.00 for discrete analysis or difference of means negative for continuous analysis.

14.4.4 Model 4: 1987 Dioxin Level Analysis

The adjusted Model 4 analysis revealed a significant positive association between essential hypertension and 1987 dioxin. A marginally significant association between the evidence of a prior myocardial infarction, as determined from the ECG, and 1987 dioxin also was observed. The results of all unadjusted and adjusted Model 4 analyses are summarized in Table 14-42.

Table 14-42. Summary of 1987 Dioxin Analysis (Model 4) for Cardiovascular Variables (Ranch Hands Only)

Variable	Unadjusted	Adjusted
Medical Records		
Essential Hypertension (D)	+<0.001	+0.011
Heart Disease (Excluding Essential Hypertension) (D)	-0.004	ns
Myocardial Infarction (D)	NS	NS
Stroke or Transient Ischemic Attack (D)	ns	NS
Physical Examination		
Systolic Blood Pressure (C)	NS	ns
Systolic Blood Pressure (D)	NS	ns*
Diastolic Blood Pressure (C)	+0.014	NS
Diastolic Blood Pressure (D)	NS	NS

Table 14-42. Summary of 1987 Dioxin Analysis (Model 4) for Cardiovascular Variables (Ranch Hands Only) (Continued)

Variable	Unadjusted	Adjusted
Heart Sounds (D)	NS	NS
Overall Electrocardiograph (ECG) (D)	ns	NS
ECG: Right Bundle Branch Block (D)	NS	NS
ECG: Left Bundle Branch Block (D)	ns	ns
ECG: Non-Specific ST- and T-Wave Changes (D)	NS	NS
ECG: Bradycardia (D)	ns*	ns
ECG: Tachycardia (D)	NS	NS
ECG: Arrhythmia (D)	ns	NS
ECG: Evidence of Prior Myocardial Infarction (D)	NS	NS*
ECG: Other Diagnoses (D)	NS	NS
Funduscopic Examination (D)	NS	NS
Carotid Bruits (D)	NS	ns
Radial Pulses (D)	ns	ns
Femoral Pulses (D)	NS	NS
Popliteal Pulses (D)	ns	NS
Dorsalis Pedis Pulses (D)	ns	NS
Posterior Tibial Pulses (D)	NS	NS
Leg Pulses (D)	NS	NS
Peripheral Pulses (D)	NS	NS
Self-reported Questionnaire		
Intermittent Claudication and Vascular Insufficiency Index (ICVI) (D)	NS	NS

NS* or ns*: Marginally significant (0.05<p≤0.10).

C: Continuous analysis.

D: Discrete analysis.

+: Relative risk ≥1.00 for discrete analysis; slope nonnegative for continuous analysis.

-: Relative risk <1.00.

P-value given if p≤0.05.

A capital "NS" denotes a relative risk of 1.00 or greater for discrete analysis or slope nonnegative for continuous analysis. A lowercase "ns" denotes a relative risk less than 1.00 for discrete analysis or slope negative for continuous analysis.

14.5 CONCLUSION

Analyses revealed that Ranch Hands had a significantly higher percentage of participants with a history of heart disease (excluding essential hypertension) than did Comparisons and, in particular, within enlisted flyers. However, the risk of disease was not significantly increased in Ranch Hand enlisted groundcrew—the military occupation with the highest dioxin levels. The association between heart disease and initial dioxin for Ranch Hands showed a negative dose-response trend, with heart disease decreasing as initial dioxin increased. Furthermore, Ranch Hands in the background and the low dioxin categories had more heart disease than did Comparisons, but this increase was not seen in Ranch Hands in the high dioxin category. Increases in tachycardia and other ECG findings, such as pre-excitation, were seen for Ranch Hands in the high dioxin category, although the analyses were based on a sparse number of abnormalities. A significant positive association between initial dioxin and evidence of prior myocardial infarction from the ECG was observed in Ranch Hands, and a marginally significant positive

association was observed between 1987 dioxin and evidence of prior myocardial infarction from the ECG. A positive association between 1987 dioxin and essential hypertension also was observed in Ranch Hands. In contrast to previous AFHS examinations, no relation was found between peripheral pulses and any measures of exposure.

In summary, in contrast to prior examinations, the current study has documented that Ranch Hands are more likely than Comparisons to have historical evidence for heart disease (excluding essential hypertension) but are no longer at greater risk for the occurrence of pulse deficits. By all other indices, the prevalence of cardiovascular disease appears similar in both cohorts. For the first time, there is evidence that levels of dioxin may be a risk factor for the development of essential hypertension and prior myocardial infarction as indicated by interpretation of the ECG. As of 1997, the verified history of essential hypertension was associated with 1987 dioxin, and the evidence of prior myocardial infarction from the ECG was associated with initial dioxin. These findings, in conjunction with the increase in the number of deaths caused by diseases of the circulatory system for Ranch Hand nonflying enlisted personnel based on the 1994 AFHS mortality update (34), showed associations with dioxin that require further observation. A biological mechanism for the relation between dioxin and heart disease is unknown at this time.

REFERENCES

- 1. Hermansky, S. J., T. L. Holcslaw, W. J. Murray, R. S. Markin, and S. J. Stohs. 1988. Biochemical and functional effects of 2,3,7,8-tetrachlorodibenzo-p-dioxin on the heart of female rats. *Toxicology and Applied Pharmacology* 95:175-84.
- 2. Canga, L., R. Levi, and A. B. Rifkind. 1988. Heart as a target organ in 2,3,7,8-tetrachlorodibenzo-p-dioxin toxicity: Decreased beta-adrenergic responsiveness and evidence of increased intracellular calcium. *Proceedings of the National Academy of Sciences* 85:905-9.
- 3. Kelling, C. K., L. A. Menahan, and R. E. Peterson. 1987. Effects of 2,3,7,8-tetrachlorodibenzo-p-dioxin treatment on mechanical function of the rat heart. *Toxicology and Applied Pharmacology* 91:497-501.
- 4. Brewster, D. W., F. Matsumura, and T. Akera. 1987. Effects of 2,3,7,8-tetrachlorodibenzo-p-dioxin on guinea pig heart muscle. *Toxicology and Applied Pharmacology* 89:408-17.
- 5. Brewster, D. W., D. W. Bommbick, and F. Matsumura. 1988. Rabbit serum hypertriglyceridemia after administration of 2,3,7,8-tetrachlorodibenzo-p-dioxin (TCDD). *Journal of Toxicology and Environmental Health* 25:495-507.
- 6. Brewster, D. W., and F. Matsumura. 1989. Differential effect of 2,3,7,8-tetrachlorodibenzo-p-dioxin on adipose tissue lipoprotein lipase activity in the guinea pig, rat, hamster, rabbit, and mink. *Comparative Biochemistry and Physiology* 93C:49-53.
- 7. Kociba, R. J., D. G. Keyes, J. E. Beyer, R. M. Carreon, C. E. Wade, D. A. Dittenber, R. P. Kalnins, L. E. Frauson, C. N. Park, S. D. Barnard, R. A. Hummel, and C. G. Humiston. 1978. Results of a two-year chronic toxicity and oncogenicity study of 2,3,7,8-tetrachlorodibenzo-p-dioxin in rats. *Toxicology and Applied Pharmacology* 46:279-303.
- 8. Cantrell, S. M., L. H. Lutz, D. E. Tillitt, and M. Hannink. 1996. Embryotoxicity of 2,3,7,8-tetrachlorodibenzo-p-dioxin (TCDD): The embryonic vasculature is a physiological target to TCDD-induced DNA damage and apoptotic cell death in Medaka (Orizias latipes). *Toxicology and Applied Pharmacology* 141:23.
- 9. Guiney, P. D., R. M. Smolowitz, R. E. Peterson, and J. J. Stegeman. 1997. Correlation of 2,3,7,8-tetrachlorodibenzo-p-dioxin induction of cytochrome P4501A in vascular endothelium with toxicity in early life stages of lake trout. *Toxicology and Applied Pharmacology* 143:256.
- 10. Schiller, C. M., C. M. Adcock, and R. A. Moore. 1985. Effect of 2,3,7,8-tetrachlorodibenzo-p-dioxin and fasting on body weight and lipid parameters in rats. *Toxicology and Applied Pharmacology* 81:356-61.
- 11. Jovanovich, L., S. Levin, and M. Kahn. 1987. Significance of mirex-caused hypoglycemia and hyperlipidemia in rats. *Journal of Biochemistry and Toxicology* 2:203-13.
- 12. Swift, L. L., T. A. Gasiewicz, and G. D. Dunn. 1981. Characterization of the hyperlipidemia in guinea pigs induced by 2,3,7,8-tetrachlorodibenzo-p-dioxin. *Toxicology and Applied Pharmacology* 59:489-99.
- 13. Moses, M., R. Lilis, K. D. Crow, J. Thornton, A. Fischbein, H. A. Anderson, and I. J. Selikoff. 1984. Health status of workers with past exposure to 2,3,7,8-tetrachlorodibenzo-p-dioxin in the manufacture of 2,4,5-trichlorophenoxyacetic acid: Comparison of findings with and without chloracne. *American Journal of Industrial Medicine* 5:161-82.

- 14. Suskind, R. R., and V. S. Hertzberg. 1984. Human health effects of 2,4,5-T and its toxic contaminants. *Journal of the American Medical Association* 251:2372-80.
- 15. Zack, J. A., and W. R. Gaffey. 1983. A mortality study of workers employed at the Monsanto company plant in Nitro, West Virginia. *Environmental Science Research* 26:575-91.
- 16. Bond, G. G., M. G. Ott, F. E. Brenner, and R. R. Cook. 1983. Medical and morbidity surveillance findings among employees potentially exposed to TCDD. *British Journal of Industrial Medicine* 40:318-24.
- 17. Von Benner, A., L. Edler, K. Mayer, and A. Zober. 1994. 'Dioxin' investigation program of the chemical industry professional association. *Arbeitsmedizin Sozialmedizin Praventivmedizin* 29:11-6.
- 18. Bertazzi, P. A., C. Zocchetti, A. C. Pesatori, S. Guercilena, M. Sanarico, and L. Radice. 1989. Mortality in an area contaminated by TCDD following an industrial incident. *Medicina del Lavoro* (Italy) 80:316-29.
- 19. Bertazzi, P. A., C. Zocchetti, A. C. Pesatori, S. Guercilena, M. Sanarico, and L. Radice. 1989. Tenyear mortality study of the population involved in the Seveso incident in 1976. *American Journal of Epidemiology* 129:1187-200.
- 20. Zack, J. A., and R. R. Suskind. 1980. The mortality experience of workers exposed to tetrachlorodibenzodioxin in a trichlorophenol process accident. *Journal of Occupational Medicine* 22:11-4.
- 21. Zober, A., M. G. Ott, and P. Messerer. 1994. Morbidity follow-up study of BASF employees exposed to 2,3,7,8-tetrachlorodibenzo-p-dioxin (TCDD) after a 1953 chemical reactor incident. *Occupational and Environmental Medicine* 51:469-86.
- 22. Ott, M. G., A. Zober, and C. Germann. 1994. Laboratory results for selected target organs in 138 individuals occupationally exposed to TCDD. *Chemosphere* 29:2423-37.
- 23. Kogan, M. D., and R. W. Clapp. 1985. Mortality among Vietnam veterans in Massachusetts, 1972-1983. Massachusetts Office of the Commissioner of Veterans Services, Agent Orange Program.
- 24. Fett, M. J., J. R. Nairn, D. M. Cobbin, and M. A. Adena. 1987. Mortality among Australian conscripts of the Vietnam conflict era. II. Causes of death. *American Journal of Epidemiology* 125:878-84.
- Anderson, H. A., L. P. Hanrahan, M. Jensen, D. Laurin, W.-Y. Yick, and P. Wiegman. 1986.
 Wisconsin Vietnam veteran mortality study: proportionate mortality ratio study results.
 Madison: Wisconsin Division of Health.
- 26. Centers for Disease Control (CDC). 1987. Postservice mortality among Vietnam veterans. *Journal of the American Medical Association* 257:790-5.
- 27. Centers for Disease Control. 1988. Health status of Vietnam veterans. II. Physical health. *Journal of the American Medical Association* 259:2708-14.
- 28. Pollei, S., F. A. Mettler Jr., C. A. Kelsey, M. R. Walters, and R. E. White. 1986. Follow-up chest radiographs in Vietnam veterans: Are they useful? *Radiology* 161:101-2.
- 29. Watanabe, K. K., H. K. Kang, and T. L. Thomas. 1991. Mortality among Vietnam veterans: with methodological considerations. *Journal of Occupational Medicine* 33:780-5.
- 30. Thomas, T. L., and H. K. Kang. 1990. Mortality and morbidity among Army Chemical Corps Vietnam veterans: a preliminary report. *American Journal of Industrial Medicine* 18:665-73.

- 31. Thomas, T. L., H. Kang, and N. Dalager. 1991. Mortality among women Vietnam veterans, 1973-1987. *American Journal of Epidemiology* 134:973-80.
- 32. Eisen, S., J. Goldberg, W. R. True, and W. G. Henderson. 1991. A co-twin control study of the effects of the Vietnam war on the self-reported physical health of veterans. *American Journal of Epidemiology* 134:49-59.
- 33. Stellman, S. D., J. M. Stellman, and J. F. Sommer, Jr. 1988. Health and reproductive outcomes among American Legionnaires in relation to combat and herbicide exposure in Vietnam. *Environmental Research* 47:150-74.
- 34. Roegner, R. H., W. D. Grubbs, M. B. Lustik, A. S. Brockman, S. C. Henderson, D. E. Williams, W. H. Wolfe, J. E. Michalek, and J. C. Miner. 1991. The Air Force Health Study: An epidemiologic investigation of health effects in Air Force personnel following exposure to herbicides. Serum dioxin analysis of 1987 examination results. NTIS: AD A 237 516-24. United States Air Force School of Aerospace Medicine, Brooks Air Force Base, Texas.
- 35. Grubbs, W. D., W. H. Wolfe, J. E. Michalek, D. E. Williams, M. B. Lustik, A. S. Brockman, S. C. Henderson, F. R. Burnett, R. G. Land, D. J. Osborne, V. K. Rocconi, M. E. Schreiber, J. C. Miner, G. L. Henriksen, and J. A. Swaby. 1995. The Air Force Health Study: An epidemiologic investigation of health effects in Air Force personnel following exposure to herbicides: Final Report. 1992 Followup Examination Results. NTIS: AD A 304 306, 304 308-316. United States Air Force School of Aerospace Medicine, Brooks Air Force Base, Texas.
- 36. Wolfe, W. H., J. E. Michalek, and J. C. Miner. 1994. The Air Force Health Study: An epidemiologic investigation of health effects in Air Force personnel following exposure to herbicides: Mortality update 1994. Epidemiologic Research Division, Armstrong Laboratory, Human Systems Center, Brooks Air Force Base, Texas.
- 37. Flesch-Janys, D., J. Berger, P. Gurn, A. Manz, S. Nagel, H. Waltsgott, and J. H. Dwyer. 1995. Exposure to polychlorinated dioxins and furans (PCDD/F) and mortality in a cohort of workers from a herbicide-producing plant in Hamburg, Federal Republic of Germany. *American Journal of Epidemiology* 142(11):1165-75.
- 38. Henriksen, G. L., N. S. Ketchum, J. E. Michalek, and J. A. Swaby. 1997. Serum dioxin and diabetes mellitus in veterans of Operation Ranch Hand. *Epidemiology* 8:252-8.
- 39. Assennato, G., D. Cervino, E. A. Emmett, G. Longo, and F. Merlo. 1989. Follow-up of subjects who developed chloracne following TCDD exposure at Seveso. *American Journal of Industrial Medicine* 16:119-25.
- 40. Fingerhut, M. A., W. E. Halperin, D. A. Marlow, L. A. Piacitelli, P. A. Honchar, M. H. Sweeney, A. L. Greife, P. A. Dill, K. Steenland, and A. J. Suruda. 1991. Cancer mortality in workers exposed to 2,3,7,8-tetrachlorodibenzo-p-dioxin. *New England Journal of Medicine* 324(4):212-8.
- 41. Martin, J. V. 1984. Lipid abnormalities in workers exposed to dioxin. *British Journal of Industrial Medicine* 41:254-6.
- 42. Ashe, W. F., and R. R. Siskind. 1982. Progress report patients from Monsanto Chemical company, Nitro, West Virginia, 1959. Quoted by Hay, AWM in: *The chemical scythe: lessons of 2,4,5-T and dioxin.* New York and London: Plenum Press.
- 43. Oliver, R. M. 1975. Toxic effects of 2,3,7,8-tetrachlorodibenzo 1,4-dioxin on laboratory workers. *British Journal of Industrial Medicine* 32:49-53.
- 44. Walker, A. E., and J. V. Martin. 1979. Lipid profiles in dioxin exposed workers. *Lancet* i:446-7.

- 45. Lathrop, G. D., W. H. Wolfe, R. A. Albanese, and P. M. Moynahan. 1984. The Air Force Health Study: An epidemiologic investigation of health effects in Air Force personnel following exposure to herbicides: Baseline Morbidity Study Results. NTIS: AD A-138-340. United States Air Force School of Aerospace Medicine, Brooks Air Force Base, Texas.
- 46. Wolfe, W. H., J. E. Michalek, J. C. Miner, A. Rahe, J. Silva, W. F. Thomas, W. D. Grubbs, M. B. Lustik, T. G. Karrison, R. H. Roegner, and D. E. Williams. 1990. Health status of Air Force veterans occupationally exposed to herbicides in Vietnam. I. Physical health. *Journal of the American Medical Association* 264:1824-31.
- 47. Hoffman, R. E., P. A. Stehr-Green, K. B. Webb, G. Evans, A. P. Knutsen, W. F. Schramm, J. L. Staake, B. B. Gibson, and K. K. Steinberg. 1986. Health effects of long-term exposure to 2,3,7,8-tetrachlorodibenzo-p-dioxin. *Journal of the American Medical Association* 255:2031-38.
- 48. Stehr, P. A., G. Stein, H. Falk, E. Sampson, S. J. Smith, K. Steinberg, K. Webb, S. Ayres, and W. Schramm. 1986. A pilot epidemiologic study of possible health effects associated with 2,3,7,8-tetrachlorodibenzo-p-dioxin contamination in Missouri. *Archives of Environmental Health* 41:16-22.
- 49. Lathrop, G. D., S. G. Machado, T. G. Karrison, W. D. Grubbs, W. F. Thomas, W. H. Wolfe, J. E. Michalek, J. C. Miner, and M. R. Peterson. 1987. The Air Force Health Study: An epidemiologic investigation of health effects in Air Force personnel following exposure to herbicides: First followup examination results. NTIS: AD A 188262. United States Air Force School of Aerospace Medicine. Brooks Air Force Base, Texas.
- 50. Knapik, J. J., A. R. L. Burse, and J. A. Vogel. 1983. Height, weight, percent body fat, and indices of adiposity for young men and women entering the Army. *Aviation, Space, and Environmental Medicine* 54:223-31.
- 51. Jenkins, C. D., R. H. Roseman, and S. J. Zyzanski. 1974. Prediction of clinical coronary heart disease by a test for the coronary-prone behavior pattern. *New England Journal of Medicine* 290(23):1271-5.
- 52. Michalek, J. E., J. L. Pirkle, S. P. Caudill, R. C. Tripathi, D. G. Patterson Jr., and L. L. Needham. 1996. Pharmacokinetics of TCDD in Veterans of Operation Ranch Hand: 10-year Followup. *Journal of Toxicology and Environmental Health* 47:209-20.
- 53. Mausner, J. S., and A. K. Bahn. 1974. *Epidemiology An Introductory Text*. Philadelphia: W. B. Saunders Company.